	58662
Access DB#	1

# SEARCH REQUEST FORM

Jan

Scientific and Technical Information Center

Requester's Full Name: Canatana Art Unit: Phone N Mail Box and Bldg/Room Location:	umber 305-433 EBD 08 Resu	Examiner #: 79271 Date: 3/16/63  7 Serial Number: 15/682743  alts Format Preferred (circle): PAPER DISK E-MAIL
If more than one search is submi	itted, please prioritiz	e searches in order of need.
•		***********
Include the elected species or structures, ke	eywords, synonyms, acron hat may have a special me	as specifically as possible the subject matter to be searched.  lyms, and registry numbers, and combine with the concept or  caning. Give examples or relevant citations, authors, etc, if  abstract.
	e see bile	
Inventors (please provide full names):	Elias Hum	berto Hermida Ochoa.
Earliest Priority Filing Date:		.1
appropriate serial number.		parent, child, divisional, or issued patent numbers) along with the
Please Se	such for	a method for treatme
of deaguagest		reulan cartilage cousa
by acted to	witic was	a method for theatmenticular carriedas a viscoelastic
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	Jan Delar Reference Lib Biotechnology & Che CM1 1E07 - 703- jan.delaval@usp	orarian mical Library
*******		
STAFF USE ONLY	Type of Search	Vendors and cost where applicable
Searcher:	NA Sequence (#)	STN
Searcher Phone #: 4498	AA Sequence (#)	Dialog
Searcher Location:	Structure (#)	Questel/Orbit
Date Searcher Picked Up: 311/03	Bibliographic V	Dr.Link
Date Completed: 3115 3	Litigation	Lexis/Nexis
Searcher Prep & Review Time:	Fulltext	Sequence Systems
Clerical Prep Time	Patent Family	WWW/Internet

Other (specify)\_

+90

#### => d his

L49

23 S L48 AND EYE?/CW

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(FILE 'HOME' ENTERED AT 16:16:14 ON 15 MAR 2003)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 16:16:25 ON 15 MAR 2003
                E CHONDROITIN SULFATE/CN
              1 S E3
L1
L2
            136 S 9007-27-6/CRN AND 7664-93-9/CRN
L3
             11 S L2 AND 2/NC
              3 S L2 AND 9067-32-7/CRN
L4
              2 S L4 NOT C4H6O2S
L5
L6
              1 S 9067-32-7
              1 S 9004-61-9
L7
              3 S 9004-61-9/CRN AND L2
rs
L9
              1 S L8 AND 3/NC
     FILE 'HCAPLUS' ENTERED AT 16:18:30 ON 15 MAR 2003
              9 S L5 OR L9
L10
           7390 S L1 OR L3
L11
          11657 S CHONDROITIN(S) (SULFATE OR SULPHATE)
L12
L13
            361 S CHONDROITINSULFATE OR CHONDROITINSULPHATE
            200 S CHONDROITIN() (SULFURIC OR SULPHURIC) () ACID
L14
          11512 S CHONDROITIN (1W) (SULFATE OR SULPHATE OR (SULFURIC OR SULPHURIC
L15
           1628 S (CHONDROITINSULFURIC OR CHONDROITINSULPHURIC) () ACID
L16
L17
          13151 S L11-L16
L18
          1406 S L6
L19
           1729 S (NA OR SODIUM) () (HYALURONATE OR HYALURON OR HYALURONIC ACID)
L20
             83 S HEALON OR HYALGAN
             25 S ARTZ OR FCH 200
L21
           1862 S L18-L21
L22
           9531 S L7
L23
L24
          12860 S HYALURONATE OR HYALURON OR HYALURONIC ACID
           2450 S HYALURONAN
L25
             65 S HYALURONAN (S) (NA OR SODIUM OR SODIUM SALT)
L26
           4638 S L17 AND L19-L26
L27
            247 S L27 AND L22
L28
     FILE 'REGISTRY' ENTERED AT 16:28:28 ON 15 MAR 2003
             11 S L1 OR L3
                SEL RN
             58 S E1-E11/CRN
L30
             56 S L30 NOT L5, L9
L31
L32
             38 S L31 NOT (MXS OR IDS)/CI
            18 S L31 NOT L32
L33
L34
            262 S CHONDROITIN(L)SULFATE
L35
             88 S L34 AND SALT
L36
             63 S L35 NOT (MXS OR IDS)/CI
L37
             31 S L36 NOT (COMPD OR WITH)
     FILE 'HCAPLUS' ENTERED AT 16:31:37 ON 15 MAR 2003
L38
            484 S L37
L39
          13232 S L17, L38
                                                                 Jan Delaval
L40
            260 S L39 AND L22
                                                              Reference Librarian
           4647 S L39 AND L23-L26
                                                          Blotechnology & Chemical Library
L42
            260 S L40, L41 AND L22
                                                            CM1 1E07 = 703-308-4498
            260 S L28, L42
L43
                                                              jan.delaval@uspto.gov
L44
             50 S L43 AND GEL?
             24 S L43 AND VISCOELAST?
L45
L46
             1 S L43 AND INTRAARTICUL?
L47
             2 S L43 AND INTRA ARTICUL?
             72 S L44, L45 NOT L46, L47
L48
```

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SEL DN AN 17
               1 S L49 AND E12-E14
L50
              49 S L48 NOT L49
L51
                 SEL DN AN 5 24 38
               3 S E15-E23 AND L51
L52
                 E CARTILAGE/CT
          11561 S E3-E25
L53
                 E E3+ALL
          14712 S E7+NT
L54
                 E JOINT/CT
L55
           4768 S E6-E28
                 E E5+ALL
           1255 S E2
L56
                 E JOINT/CT
                 E E6+ALL
           8912 S E6, E5+NT
L57
L58
           2604 S E13+NT
                 E OSTEOARTHRITIS/CT
L59
           1853 S E3
                 E E3+ALL
           2870 S E11, E12, E10+NT
L60
L61
              7 S CHONDRAL (L) LESION
L62
              72 S ?CHONDRAL? (L) LESION
L63
              17 S L43 AND L53-L62
                 SEL DN AN 3
L64
               1 S L63 AND E1-E3
                 SEL DN AN L63 1 5 17
L65
               3 S E4-E12 AND L63
L66
              15 S L10, L50, L52, L64, L65
                 E OCHOA/AU
               7 S E95
L67
                 E HERMIDA/AU
                 E HUMBERTO/AU
                 E ALCON/PA, CS
                 E ALCOM/PA,CS
L68
             786 S E3-E8
             785 S ALCON?/PA,CS
             12 S L67-L69 AND L43
L71
               1 S L67-L69 AND L10
L72
              2 S L67-L69 AND L66
L73
             15 S L66, L71, L72
L74
             10 S L70 NOT L73
             15 S L73 AND L10-L28, L38-L74
L75
L76
              10 S L74 AND L10-L28, L38-L75
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#### => fil reg

FILE 'REGISTRY' ENTERED AT 16:53:16 ON 15 MAR 2003 USE IS SUBJECT TO THE TERMS OF YOUR STN CUSTOMER AGREEMENT. PLEASE SEE "HELP USAGETERMS" FOR DETAILS. COPYRIGHT (C) 2003 American Chemical Society (ACS)

Property values tagged with IC are from the ZIC/VINITI data file provided by InfoChem.

STRUCTURE FILE UPDATES: 13 MAR 2003 HIGHEST RN 499099-49-9 DICTIONARY FILE UPDATES: 13 MAR 2003 HIGHEST RN 499099-49-9

TSCA INFORMATION NOW CURRENT THROUGH MAY 20, 2002

Please note that search-term pricing does apply when conducting SmartSELECT searches.

Crossover limits have been increased. See HELP CROSSOVER for details.

Experimental and calculated property data are now available. See HELP PROPERTIES for more information. See STNote 27, Searching Properties in the CAS Registry File, for complete details: http://www.cas.org/ONLINE/STN/STNOTES/stnotes27.pdf

=> s 15 or 19 3 L5 OR L9 L77

=> d ide can tot

L77 ANSWER 1 OF 3 REGISTRY COPYRIGHT 2003 ACS

153311-76-3 REGISTRY RN

Chondroitin, hydrogen sulfate, mixt. with hyaluronic acid sodium salt CN (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

Hyaluronic acid, sodium salt, mixt. contg. (9CI) CN

H2 O4 S . x Unspecified . Unspecified MF

CI MXS

SR CA

LCSTN Files: CA, CAPLUS, USPATFULL

> CM 1

CRN 9067-32-7

CMF Unspecified

CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

2 CM

CRN 9007-28-7

H2 O4 S . x Unspecified CMF

> CM 3

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4

7664-93-9 CRN

CMF H2 O4 S

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 120:144233

L77 ANSWER 2 OF 3 REGISTRY COPYRIGHT 2003 ACS

123352-36-3 REGISTRY RN

李孝子 李素素 不 一

CN Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid sodium salt (9CI) (CA INDEX NAME)

OTHER CA INDEX NAMES:

CN Hyaluronic acid, sodium salt, mixt. contg. (9CI)

OTHER NAMES:

CN Viscoat

MF H2 O4 S . x Na . x Unspecified . Unspecified

CI MXS

SR CA

LC STN Files: BIOBUSINESS, BIOSIS, CA, CAPLUS, CIN, MEDLINE, PHARMASEARCH, PROMT, TOXCENTER, USPATFULL

CM 1

CRN 9067-32-7

CMF Unspecified

CCI PMS, MAN

#### \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 9082-07-9

CMF  $\mbox{H2 O4 S}$  .  $\mbox{x Na}$  .  $\mbox{x Unspecified}$ 

CM 3

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4

CRN 7664-93-9

CMF H2 O4 S

- 7 REFERENCES IN FILE CA (1962 TO DATE)
- 7 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:83679

REFERENCE 2: 135:55996

REFERENCE 3: 131:139516

REFERENCE 4: 130:32985

REFERENCE 5: 127:824

REFERENCE 6: 122:274105

REFERENCE 7: 111:187541

```
L77 ANSWER 3 OF 3 REGISTRY COPYRIGHT 2003 ACS
     108145-77-3 REGISTRY
RN
     Chondroitin, hydrogen sulfate, mixt. with hyaluronic acid (9CI) (CA INDEX
CN
     NAME)
OTHER CA INDEX NAMES:
     Hyaluronic acid, mixt. contg. (9CI)
CN
     H2 O4 S . x Unspecified . Unspecified
MF
CI
     MXS
SR
     CA
LC
     STN Files: CA, CAPLUS
     CM
          1
         9004-61-9
     CRN
         Unspecified
     CMF
     CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN 9007-28-7
     CMF
         H2 O4 S . x Unspecified
               3
          CM
          CRN
               9007-27-6
               Unspecified
          CMF
               PMS, MAN
          CCI
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          CM
               4
          CRN
              7664-93-9
          CMF H2 O4 S
   0
HO-S-OH
   0
               1 REFERENCES IN FILE CA (1962 TO DATE)
               1 REFERENCES IN FILE CAPLUS (1962 TO DATE)
            1: 106:188973
REFERENCE
=> d ide can 16
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
L6
     9067-32-7 REGISTRY
RN
     Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
OTHER NAMES:
CN
     Artz
CN
     Bio Hyaluro 12
CN
     FCH 200
CN
     FCH 248
```

CN

CN

HA-O

HA-Q 1

```
CN
     Healon
     Healon (polysaccharide)
CN
     Healon GV
CN
     Hyalart
CN
     Hyalein
CN
CN
     Hyalgan
CN
     Hyladerm
CN
     Nidelon
CN
     NRD 101
CN
     Opegan
CN
     Orthovisc
CN
     SI 4402
     SL 1010
CN
CN
     SLM 10
CN
     Sodium hyaluronate
CN
     SPH
DR
     34448-35-6
MF
     Unspecified
CI
     PMS, COM, MAN
     Manual registration, Polyother, Polyother only
PCT
                  ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
LC
       BIOSIS, BIOTECHNO, CA, CAPLUS, CASREACT, CBNB, CHEMCATS, CHEMLIST, CIN,
       CSCHEM, DDFU, DIOGENES, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA,
       MRCK*, PHAR, PHARMASEARCH, PROMT, RTECS*, TOXCENTER, USAN, USPAT2,
       USPATFULL
         (*File contains numerically searchable property data)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
            1402 REFERENCES IN FILE CA (1962 TO DATE)
              57 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            1406 REFERENCES IN FILE CAPLUS (1962 TO DATE)
            1: 138:158867
REFERENCE
REFERENCE
            2:
                138:158860
                138:158840
REFERENCE
            3:
REFERENCE
            4:
                138:142492
REFERENCE
            5:
                138:127015
REFERENCE
            6:
                138:126891
                138:117178
REFERENCE
            7:
                138:112513
REFERENCE
            8:
                138:112480
REFERENCE
            9:
REFERENCE 10: 138:112252
=> d ide can 17
     ANSWER 1 OF 1 REGISTRY COPYRIGHT 2003 ACS
1.7
RN
     9004-61-9 REGISTRY
CN
     Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
OTHER NAMES:
CN
     ACP
CN
     ACP (polysaccharide)
CN
     ACP gel
```

CN

Durolane

```
Hyaluronan
CN
     Hylartil
CN
CN
     Luronit
     Mucoitin
CN
CN
     Sepracoat
CN
     Sofast
CN
     Synvisc
DR
     9039-38-7, 37243-73-5, 29382-75-0
MF
     Unspecified
CI
     PMS, COM, MAN
PCT
     Manual registration, Polyester, Polyester formed
     STN Files: ADISINSIGHT, ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS,
LC
       BIOSIS, BIOTECHNO, CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN,
       CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DIOGENES, DRUGNL, DRUGU,
       DRUGUPDATES, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK*,
       NAPRALERT, NIOSHTIC, PHAR, PHARMASEARCH, PIRA, PROMT, TOXCENTER, USAN,
       USPAT2, USPATFULL
         (*File contains numerically searchable property data)
                      DSL**, EINECS**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
            9448 REFERENCES IN FILE CA (1962 TO DATE)
            711 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            9462 REFERENCES IN FILE CAPLUS (1962 TO DATE)
            1: 138:175923
REFERENCE
REFERENCE
            2: 138:175749
REFERENCE
            3: 138:175701
REFERENCE
            4: 138:175682
            5: 138:172181
REFERENCE
REFERENCE
            6: 138:170404
REFERENCE
            7: 138:168381
REFERENCE
            8:
               138:168132
REFERENCE
            9:
               138:167834
REFERENCE 10: 138:167658
=> d ide can tot 129
L29 ANSWER 1 OF 11 REGISTRY COPYRIGHT 2003 ACS
     185323-66-4 REGISTRY
RN
CN
     Chondroitin, octakis(hydrogen sulfate) (9CI) (CA INDEX NAME)
MF
     H2 O4 S . 1/8 Unspecified
SR
     CA
     STN Files: CA, CAPLUS, TOXCENTER, USPATFULL
LC
     CM
          9007-27-6
     CRN
     CMF
          Unspecified
          PMS, MAN
     CCI
```

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 126:141392

REFERENCE 2: 126:75185

L29 ANSWER 2 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 163063-19-2 REGISTRY

CN Chondroitin, tris(hydrogen sulfate) (ester) (9CI) (CA INDEX NAME)

MF H2 O4 S . 1/3 Unspecified

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

1 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 122:289056

L29 ANSWER 3 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 143928-11-4 REGISTRY

CN Chondroitin, tetrakis(hydrogen sulfate) (ester) (9CI) (CA INDEX NAME)

MF H2 O4 S . 1/4 Unspecified

PCT Manual registration

SR CA

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

CM 1

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

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3 REFERENCES IN FILE CA (1962 TO DATE)

3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 126:141392

REFERENCE 2: 126:75185

REFERENCE 3: 117:178119

L29 ANSWER 4 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 143928-10-3 REGISTRY

CN Chondroitin, hexakis(hydrogen sulfate) (ester) (9CI) (CA INDEX NAME)

MF H2 O4 S . 1/6 Unspecified

PCT Manual registration

SR CF

LC STN Files: CA, CAPLUS, TOXCENTER, USPATFULL

CM 1

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 126:75185

REFERENCE 2: 117:178119

L29 ANSWER 5 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 129837-56-5 REGISTRY

CN Chondroitin, 2'-(hydrogen sulfate) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Chondroitin 8-sulfate

MF H2 O4 S . Unspecified

CI COM

PCT Manual registration

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

# \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9

CMF H2 O4 S

- 3 REFERENCES IN FILE CA (1962 TO DATE)
- 3 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 130:100352

REFERENCE 2: 115:64715

REFERENCE 3: 113:154827

L29 ANSWER 6 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 129837-55-4 REGISTRY

CN Chondroitin, 2',4-bis(hydrogen sulfate) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Chondroitin 4,8-disulfate

MF H2 O4 S . 1/2 Unspecified

PCT Manual registration

SR CA

LC STN Files: CA, CAPLUS

CM 1

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

# \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

2 REFERENCES IN FILE CA (1962 TO DATE)

2 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 130:100352

REFERENCE 2: 113:154827

L29 ANSWER 7 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 81988-93-4 REGISTRY

CN Chondroitin, 4,6-bis(hydrogen sulfate) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Chondroitin 4,6-disulfate

CN Chondroitin 4/6-sulfate

CN Chondroitin sulfate A-chondroitin sulfate C mixture

CN Chondroitin sulfate AC

CN Chondroitinsulfuric acid, type AC

DR 58449-35-7

MF H2 O4 S . 1/2 Unspecified

CI COM

PCT Manual registration

LC STN Files: BIOSIS, CA, CAPLUS, TOXCENTER, USPATFULL

CM 1

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

#### \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9

CMF H2 O4 S

61 REFERENCES IN FILE CA (1962 TO DATE)

1 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA

61 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 136:236879

REFERENCE 2: 135:17239

REFERENCE 3: 134:190011

REFERENCE 4: 132:178219

REFERENCE 5: 129:160200

REFERENCE 6: 129:50629

REFERENCE 7: 128:189802

REFERENCE 8: 128:100690

REFERENCE 9: 128:99486

REFERENCE 10: 127:203952

L29 ANSWER 8 OF 11 REGISTRY COPYRIGHT 2003 ACS

RN 50814-15-8 REGISTRY

CN Chondroitin, 2',6-bis(hydrogen sulfate) (9CI) (CA INDEX NAME)

OTHER NAMES:

CN Chondroitin 2',6-disulfate

CN Chondroitin, sulfate D

CN Chondroitinsulfuric acid, type D

MF H2 O4 S . 1/2 Unspecified

PCT Manual registration

LC STN Files: ANABSTR, BIOSIS, CA, CAPLUS, CHEMCATS, MEDLINE, TOXCENTER

CM 1

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

### \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9

CMF H2 O4 S

- 37 REFERENCES IN FILE CA (1962 TO DATE)
- 2 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
- 37 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 137:120035

REFERENCE 2: 136:319391

REFERENCE 3: 135:339800

REFERENCE 4: 135:147692

REFERENCE 5: 134:37548

REFERENCE 6: 133:322084

REFERENCE 7: 133:160235 8: 132:292466 REFERENCE 9: 132:104396 REFERENCE REFERENCE 10: 131:68233 L29 ANSWER 9 OF 11 REGISTRY COPYRIGHT 2003 ACS 25322-46-7 REGISTRY RN CN Chondroitin, 6-(hydrogen sulfate) (9CI) (CA INDEX NAME) OTHER CA INDEX NAMES: Chondroitinsulfuric acids, type C (8CI) OTHER NAMES: CN Chondroitin 6-sulfate CN Chondroitin C sulfate CN Chondroitin sulfate C CN Chondroitin sulfate type C CN Chondroitin sulfuric acid C CN Chondroitin sulphate C CN Chondroitin-6-sulfuric acid CN Chondroitinsulfuric acid, type C DR 9045-60-7, 49718-76-5 MF H2 O4 S . Unspecified CI STN Files: AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO, CA, LC CANCERLIT, CAPLUS, CASREACT, CHEMCATS, CHEMLIST, CSCHEM, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE, MRCK\*, PROMT, TOXCENTER, USPAT2, USPATFULL (\*File contains numerically searchable property data) EINECS\*\* Other Sources: (\*\*Enter CHEMLIST File for up-to-date regulatory information) CM1 CRN 9007-27-6 CMF Unspecified CCI PMS, MAN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* CM2 CRN 7664-93-9 CMF H2 O4 S 0 HO- S- OH 1872 REFERENCES IN FILE CA (1962 TO DATE) 97 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA 1873 REFERENCES IN FILE CAPLUS (1962 TO DATE) 1: 138:158745 REFERENCE

REFERENCE

REFERENCE

2: 138:126851

3: 138:102281

```
4: 138:100348
REFERENCE
            5: 138:95059
REFERENCE
REFERENCE
            6: 138:88443
REFERENCE
            7: 138:87021
REFERENCE
            8:
               138:83028
REFERENCE
            9: 138:70386
REFERENCE 10: 138:44720
L29 ANSWER 10 OF 11 REGISTRY COPYRIGHT 2003 ACS
     24967-93-9 REGISTRY
     Chondroitin, 4-(hydrogen sulfate) (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Chondroitinsulfuric acids, type A (8CI)
OTHER NAMES:
CN
     Chondranol
     Chondroitin 4-sulfate
CN
     Chondroitin 4-sulfuric acid
CN
CN
     Chondroitin A sulfate
CN
    Chondroitin sulfate A
CN
    Chondroitin sulfate type A
CN
     Chondroitinsulfuric acid A
CN
     Chondroitinsulfuric acid type A
     Chondroitinsulfuric acid, type A
CN
     Org 10172
CN
     Translagen
CN
CN
     Turkadon
     12643-04-8, 9040-92-0, 9045-58-3
DR
     H2 O4 S . Unspecified
MF
CI
     COM
PCT
     Manual registration
     STN Files: ADISINSIGHT, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS,
LC
       BIOTECHNO, CA, CAPLUS, CHEMCATS, CHEMLIST, CIN, CSCHEM, DDFU, DRUGU,
       EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MRCK*, NAPRALERT, PROMT, RTECS*,
       TOXCENTER, USAN, USPAT2, USPATFULL, VETU
         (*File contains numerically searchable property data)
                      EINECS**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
     CM
          1
          9007-27-6
     CRN
     CMF
          Unspecified
     CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
     CRN 7664-93-9
     CMF H2 O4 S
```

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о
но-s-он
|
о
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2165 REFERENCES IN FILE CA (1962 TO DATE)
              96 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
            2167 REFERENCES IN FILE CAPLUS (1962 TO DATE)
            1: 138:102281
REFERENCE
REFERENCE
            2:
               138:100348
               138:88443
REFERENCE
            3:
REFERENCE
               138:88197
            4:
                138:70386
REFERENCE
            5:
                138:63051
REFERENCE
            6:
REFERENCE
            7:
               138:44720
REFERENCE
            8:
                138:33368
            9:
                138:22339
REFERENCE
                137:380014
REFERENCE
          10:
L29 ANSWER 11 OF 11 REGISTRY COPYRIGHT 2003 ACS
     9007-28-7 REGISTRY
     Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
OTHER CA INDEX NAMES:
     Chondroitinsulfuric acids (8CI)
CN
OTHER NAMES:
CN
     Chondroitin polysulfate
CN
     Chondroitin sulfate
     Chondroitin sulphate
CN
     Chondroitinsulfuric acid
CN
CN
     Chonsurid
     9046-20-2, 9062-29-7, 11120-14-2, 56480-79-6
DR
MF
     H2 O4 S . x Unspecified
CI
     COM
PCT Manual registration
LC
     STN Files: ADISNEWS, AGRICOLA, ANABSTR, BIOBUSINESS, BIOSIS, BIOTECHNO,
       CA, CABA, CANCERLIT, CAPLUS, CASREACT, CBNB, CEN, CHEMCATS, CHEMLIST,
       CIN, CSCHEM, DDFU, DRUGU, EMBASE, IFICDB, IFIPAT, IFIUDB, IPA, MEDLINE,
       MRCK*, NAPRALERT, NIOSHTIC, PHAR, PROMT, RTECS*, TOXCENTER, USPAT2,
       USPATFULL
         (*File contains numerically searchable property data)
                    EINECS**, NDSL**, TSCA**
     Other Sources:
         (**Enter CHEMLIST File for up-to-date regulatory information)
     CM
          1
          9007-27-6
     CRN
     CMF
          Unspecified
     CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
```

CM 2

CRN 7664-93-9 CMF H2 O4 S

4890 REFERENCES IN FILE CA (1962 TO DATE)
314 REFERENCES TO NON-SPECIFIC DERIVATIVES IN FILE CA
4896 REFERENCES IN FILE CAPLUS (1962 TO DATE)

REFERENCE 1: 138:175914

REFERENCE 2: 138:172181

REFERENCE 3: 138:168132

REFERENCE 4: 138:168094

REFERENCE 5: 138:166260

REFERENCE 6: 138:166242

REFERENCE 7: 138:158905

REFERENCE 8: 138:158821

REFERENCE 9: 138:150822

REFERENCE 10: 138:142516

#### => fil hcaplus

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FILE COVERS 1907 - 15 Mar 2003 VOL 138 ISS 12 FILE LAST UPDATED: 14 Mar 2003 (20030314/ED)

This file contains CAS Registry Numbers for easy and accurate substance identification.

#### => d all hitstr tot 175

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ANSWER 1 OF 15 HCAPLUS COPYRIGHT 2003 ACS
L75
     2003:76549 HCAPLUS
ΑN
DN
    138:112513
    Cartilage repair and regeneration scaffold and method
TI
     Plouhar, Pamela Lynn; Malaviya, Prasanna; Schwartz, Herbert Eugene
ΙN
     Depuy Products, Inc., USA
PA
     PCT Int. Appl., 31 pp.
SO
    CODEN: PIXXD2
DT
    Patent
    English
LA
IC
    ICM A61K
     63-7 (Pharmaceuticals)
CC
     Section cross-reference(s): 9
FAN.CNT 11
                                           APPLICATION NO. DATE
     PATENT NO.
                      KIND DATE
                     A2 20030130 WO 2002-US22411 20020715
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    WO 2003007879
PΙ
         W: AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
             CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
             GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
             LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
             PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ, UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
             TJ, TM
         RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, BG,
             CH, CY, CZ, DE, DK, EE, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, SK, TR, BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR,
             NE, SN, TD, TG
                                           US 2002-195334
     US 2003033021
                     A1
                            20030213
                                                             20020715
                     Ρ
                            20010716
PRAI US 2001-305786P
                      Ρ
     US 2002-388724P
                            20020614
    A method for the repair of a cartilaginous tissue defect and a cartilage
AΒ
    repair device are disclosed. In the method for the repair of a
     cartilaginous tissue defect, a device comprising a synthetic polymer is
     implanted into a space subsequent to removal of the defect, and a biol.
     lubricant is administered at the site of the defect. The device comprises
     a synthetic polymer and a biol. lubricant.
ST
     cartilage repair scaffold polymer lubricant
TT
        (basement membrane; cartilage repair and regeneration scaffold and
        method)
IT
     Lubricants
        (biol.; cartilage repair and regeneration scaffold and method)
ΙT
     Animal tissue culture
       Cartilage
     Extracellular matrix
     Regeneration, animal
     Sterilization and Disinfection
     Surgery
        (cartilage repair and regeneration scaffold and method)
     Polymers, biological studies
IT
     RL: DEV (Device component use); TEM (Technical or engineered material
     use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cartilage repair and regeneration scaffold and method)
ΙT
     Head
        (comb, hyaluronate lubricant from; cartilage repair and
        regeneration scaffold and method)
     Prosthetic materials and Prosthetics
IT
        (implants; cartilage repair and regeneration scaffold and method)
IT
     Joint, anatomical
```

(knee, repair of; cartilage repair and regeneration scaffold

```
and method)
    Synovial fluid
ΙT
        (lubricant; cartilage repair and regeneration scaffold and method)
IT
    Glycosaminoglycans, biological studies
    Vitronectin
    RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lubricant; cartilage repair and regeneration scaffold and method)
IT
    Textiles
        (medical; cartilage repair and regeneration scaffold and method)
ΙT
    Joint, anatomical
        (meniscus; cartilage repair and regeneration scaffold and
       method)
ΙT
    Intestine
        (small, submucosa; cartilage repair and regeneration scaffold and
       method)
IT
    Bladder
    Stomach
        (submucosa; cartilage repair and regeneration scaffold and method)
     2453-03-4, Trimethylene carbonate 9002-89-5, Polyvinyl alcohol
IT
    24980-41-4, Polycaprolactone 25248-42-4, Polycaprolactone
                                                                   26009-03-0,
    Polyglycolic acid 26023-30-3, Poly[oxy(1-methyl-2-oxo-1,2-ethanediyl)]
    26100-51-6, Polylactic acid 26124-68-5, Polyglycolic acid
                                                                   31621-87-1,
    Polydioxanone
    RL: DEV (Device component use); TEM (Technical or engineered material
    use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cartilage repair and regeneration scaffold and method)
    9004-61-9, Hyaluronic acid 9007-28-7
ΤТ
                             9050-30-0, Heparan sulfate
     , Chondroitin sulfate
    9056-36-4, Keratan sulfate 9067-32-7, Sodium
                   24967-94-0, Dermatan sulfate
    hyaluronate
    RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lubricant; cartilage repair and regeneration scaffold and method)
    9004-61-9, Hyaluronic acid 9007-28-7
IT
     , Chondroitin sulfate 9067-32-7,
    Sodium hyaluronate
    RL: BUU (Biological use, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (lubricant; cartilage repair and regeneration scaffold and method)
     9004-61-9 HCAPLUS
RN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9007-28-7 HCAPLUS
RN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
         1
         9007-27-6
    CRN
         Unspecified
    CMF
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
         2
    CRN 7664-93-9
     CMF H2 O4 S
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O
||
HO-S-OH
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RN 9067-32-7 HCAPLUS

CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

L75 ANSWER 2 OF 15 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:695831 HCAPLUS

DN **137:237785** 

TI Porous beta-tricalcium phosphate granules for bone implantation, and methods for producing same

IN Dalal, Paresh S.; Dimaano, Godofredo R.; Toth, Carol Ann; Kulkarni, Shailesh C.

PA Stryker Corporation, USA

SO PCT Int. Appl., 151 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61L027-12 ICS A61L027-56

CC 63-7 (Pharmaceuticals)

Section cross-reference(s): 2, 15

FAN.CNT 1

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APPLICATION NO.
                                                         DATE
    PATENT NO.
                    KIND DATE
                    ____
                                        _____
    _____
                                                        _____
                   A2
ΡI
    WO 2002070029
                          20020912
                                         WO 2002-US5827
                                                         20020226
    WO 2002070029
                    A3
                          20030206
           AE, AG, AL, AM, AT, AU, AZ, BA, BB, BG, BR, BY, BZ, CA, CH, CN,
            CO, CR, CU, CZ, DE, DK, DM, DZ, EC, EE, ES, FI, GB, GD, GE, GH,
            GM, HR, HU, ID, IL, IN, IS, JP, KE, KG, KP, KR, KZ, LC, LK, LR,
            LS, LT, LU, LV, MA, MD, MG, MK, MN, MW, MX, MZ, NO, NZ, OM, PH,
            PL, PT, RO, RU, SD, SE, SG, SI, SK, SL, TJ, TM, TN, TR, TT, TZ,
            UA, UG, US, UZ, VN, YU, ZA, ZM, ZW, AM, AZ, BY, KG, KZ, MD, RU,
            TJ, TM
        RW: GH, GM, KE, LS, MW, MZ, SD, SL, SZ, TZ, UG, ZM, ZW, AT, BE, CH,
            CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE, TR,
            BF, BJ, CF, CG, CI, CM, GA, GN, GQ, GW, ML, MR, NE, SN, TD, TG
                                       US 2001-798518 20010302
    US 2003049328
                    A1
                          20030313
PRAI US 2001-798518
                     Α
                          20010302
    US 2001-960789
                     Α
                          20010921
```

- AB A porous .beta.-tricalcium phosphate material for bone implantation is provided. The multiple pores in the porous TCP body are sep. discrete voids and are not interconnected. The pore size diam. is in the range of 20-500 .mu.m, preferably 50-125 .mu.m. The porous .beta.-TCP material provides a carrier matrix for bioactive agents and can form a moldable putty compn. upon the addn. of a binder. Preferably, the bioactive agent is encapsulated in a biodegradable agent. The invention provides a kit and an implant device comprising the porous .beta.-TCP, and a bioactive agent and a binder. The invention also provides an implementable prosthetic device comprising a prosthetic implant having a surface region, a porous .beta.-TCP material disposed on the surface region optionally comprising at least a bioactive agent or a binder. Methods of producing the porous .beta.-TCP material and including bone formation are also provided.
- ST bone implant porous beta tricalcium phosphate granule sequence
- IT Bone morphogenetic proteins

IT

TΤ

TТ

IT

TT

TΨ

ΙT

ΙT

IT

IT

IT

ΙT

IT

implantation)

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RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (2; porous .beta.-tricalcium phosphate granules for bone implantation)
Bone morphogenetic proteins
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (3; porous .beta.-tricalcium phosphate granules for bone implantation)
Bone morphogenetic proteins
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (4; porous .beta.-tricalcium phosphate granules for bone implantation)
Bone morphogenetic proteins
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (5; porous .beta.-tricalcium phosphate granules for bone implantation)
Bone morphogenetic proteins
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (6; porous .beta.-tricalcium phosphate granules for bone implantation)
Bone morphogenetic proteins
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (7; porous .beta.-tricalcium phosphate granules for bone implantation)
Nucleic acids
RL: PEP (Physical, engineering or chemical process); PYP (Physical
process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
USES (Uses)
   (BMP-encoding; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Carbohydrates, biological studies
RL: TEM (Technical or engineered material use); THU (Therapeutic use);
BIOL (Biological study); USES (Uses)
   (aldonic acids, polymer; porous .beta.-tricalcium phosphate granules
   for bone implantation)
Transplant and Transplantation
   (allotransplant; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Polyesters, biological studies
RL: TEM (Technical or engineered material use); THU (Therapeutic use);
BIOL (Biological study); USES (Uses)
   (arom.; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Bone
  Hip
   (artificial; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Transplant and Transplantation
   (autotransplant; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Polymers, biological studies
RL: TEM (Technical or engineered material use); THU (Therapeutic use);
BIOL (Biological study); USES (Uses)
   (biodegradable; porous .beta.-tricalcium phosphate granules for bone
   implantation)
Glues
   (fibrin-contg.; porous .beta.-tricalcium phosphate granules for bone
```

```
ΙT
    Drug delivery systems
        (granules; porous .beta.-tricalcium phosphate granules for bone
        implantation)
IΤ
     Drug delivery systems
    Prosthetic materials and Prosthetics
        (implants; porous .beta.-tricalcium phosphate granules for bone
        implantation)
ΙT
    Putty
        (medical; porous .beta.-tricalcium phosphate granules for bone
        implantation)
TΤ
    Polyethers, biological studies
    RL: TEM (Technical or engineered material use); THU (Therapeutic use);
    BIOL (Biological study); USES (Uses)
        (ortho ester group-contg.; porous .beta.-tricalcium phosphate granules
        for bone implantation)
ΙT
    Growth factors, animal
    RL: PEP (Physical, engineering or chemical process); PYP (Physical
    process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
    USES (Uses)
        (osteogenins; porous .beta.-tricalcium phosphate granules for bone
        implantation)
ΙT
    Polyimides, biological studies
    RL: TEM (Technical or engineered material use); THU (Therapeutic use);
    BIOL (Biological study); USES (Uses)
        (polyanhydride-; porous .beta.-tricalcium phosphate granules for bone
        implantation)
ΙT
    Polyanhydrides
    RL: TEM (Technical or engineered material use); THU (Therapeutic use);
    BIOL (Biological study); USES (Uses)
        (polyimide-; porous .beta.-tricalcium phosphate granules for bone
        implantation)
ΙT
    Binders
    Encapsulation
    Granulation
    Mammalia
    Molecular weight distribution
    Particle size distribution
    Porosity
    Prosthetic materials and Prosthetics
    Protein sequences
    Sieving
    Sintering
    Sublimation
    cDNA sequences
        (porous .beta.-tricalcium phosphate granules for bone implantation)
    Gelatins, biological studies
IT
    Glycosaminoglycans, biological studies
    Mucins
    Peptides, biological studies
    Petrolatum
    Polyamides, biological studies
    Polyoxyalkylenes, biological studies
    Polysaccharides, biological studies
     Polyurethanes, biological studies
    RL: MOA (Modifier or additive use); TEM (Technical or engineered material
    use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
ΙT
     Polyvinyl butyrals
    RL: NUU (Other use, unclassified); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
ΙT
     Bone morphogenetic proteins
     RL: PEP (Physical, engineering or chemical process); PRP (Properties); PYP
     (Physical process); THU (Therapeutic use); BIOL (Biological study); PROC
```

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(Process); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
IΤ
     Interleukin 6
    RL: PEP (Physical, engineering or chemical process); PYP (Physical
    process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
    USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
ΙT
    Collagens, biological studies
     Polyanhydrides
    Polyphosphazenes
    RL: TEM (Technical or engineered material use); THU (Therapeutic use);
    BIOL (Biological study); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
TΤ
    Drug delivery systems
        (powders; porous .beta.-tricalcium phosphate granules for bone
        implantation)
     Fats and Glyceridic oils, biological studies
TΤ
    RL: MOA (Modifier or additive use); TEM (Technical or engineered material
    use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (sesame; porous .beta.-tricalcium phosphate granules for bone
        implantation)
ΙT
    Drug delivery systems
        (sustained-release; porous .beta.-tricalcium phosphate granules for
       bone implantation)
IT
    Transforming growth factors
    RL: PEP (Physical, engineering or chemical process); PYP (Physical
    process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
    USES (Uses)
        (.beta.-; porous .beta.-tricalcium phosphate granules for bone
        implantation)
IT
     9001-78-9, Alkaline phosphatase
    RL: BSU (Biological study, unclassified); BUU (Biological use,
    unclassified); BIOL (Biological study); USES (Uses)
        (osteogenesis marker; porous .beta.-tricalcium phosphate granules for
       bone implantation)
    7758-87-4, .beta.-Tricalcium phosphate
ΙT
    RL: DEV (Device component use); PRP (Properties); TEM (Technical or
     engineered material use); THU (Therapeutic use); BIOL (Biological study);
    USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
                        9004-54-0, Dextran, biological studies
TΤ
     69-65-8, Mannitol
    9004-61-9, Hyaluronic acid
                                  9004-62-0,
    Hydroxyethylcellulose 9004-65-3, Hydroxypropyl methylcellulose
    9005-38-3, Sodium alginate 9007-28-7, Chondroitin
               9012-76-4, Chitosan
                                     9032-42-2, Hydroxyethyl
    methylcellulose
                       9041-56-9, Hydroxybutyl methylcellulose
                                                                 9050-04-8
    9067-32-7, Sodium hyaluronate
                                     9078-35-7
                  25322-68-3, Polyethylene glycol
                                                    25513-46-6, Polyglutamic
    24991-23-9
                                          26124-68-5, Polyglycolic acid
    acid
            26100-51-6, Polylactic acid
                  52352-27-9, Polyhydroxybutyric acid
                                                        78644-42-5, Polymalic
    34346-01-5
    acid
            106392-12-5, Poloxamer
    RL: MOA (Modifier or additive use); TEM (Technical or engineered material
    use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
     9002-89-5, Polyvinyl alcohol
                                    9003-39-8, Polyvinylpyrrolidone
TT
     9004-36-8, Cellulose acetate butyrate
                                             9005-25-8, Starch, uses
    RL: NUU (Other use, unclassified); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
                               50-28-2, Estradiol, biological studies
IT
     50-23-7, Hydrocortisone
     57-83-0, Progesterone, biological studies
                                                 302-79-4, Retinoic acid
                            9002-64-6, Pth 9002-72-6, Growth hormone
     1406-16-2, Vitamin d
     9004-10-8, Insulin, biological studies 62031-54-3, Fgf
                                                                67763-96-6,
     Igf-i
```

```
RL: PEP (Physical, engineering or chemical process); PYP (Physical
    process); THU (Therapeutic use); BIOL (Biological study); PROC (Process);
    USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
IΤ
     9003-01-4D, Polyacrylic acid, derivs.
                                            24937-78-8, Ethylene-vinyl acetate
               24980-41-4, Poly(caprolactone) 25248-42-4,
                                                      26023-30-3,
                         26009-03-0, Polyglycolide
    Poly(caprolactone)
    Poly[oxy(1-methyl-2-oxo-1,2-ethanediyl)] 26063-00-3, Polyhydroxybutyrate
                 26202-08-4, Polyglycolide 26680-10-4, Poly(D,L-lactide)
     26161-42-2
    26744-04-7
                 26780-50-7, Polyglactin
                                          29223-92-5, Poly(p-dioxanone)
                                               33135-50-1, Poly(L-lactide)
     31852-84-3, Poly(trimethylene carbonate)
     41706-81-4, Poly(..epsilon..-caprolactone-glycolide) 50862-75-4,
                                           75734-93-9, Poly(glycolide-
    Poly(oxycarbonyloxy-1,3-propanediyl)
                             129515-24-8, Poly(D,L-lactide-trimethylene
    trimethylene carbonate)
     carbonate)
    RL: TEM (Technical or engineered material use); THU (Therapeutic use);
    BIOL (Biological study); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
     458061-50-2
IT
    RL: PRP (Properties)
        (unclaimed nucleotide sequence; porous beta-tricalcium phosphate
       granules for bone implantation, and methods for producing same)
                                               458061-44-4
                                                             458061-45-5
                   458061-42-2
                                 458061-43-3
ΙT
     458061-41-1
                   458061-47-7
                                 458061-48-8
                                               458061-49-9
     458061-46-6
    RL: PRP (Properties)
        (unclaimed protein sequence; porous beta-tricalcium phosphate granules
        for bone implantation, and methods for producing same)
    9004-61-9, Hyaluronic acid 9007-28-7
TT
     , Chondroitin sulfate 9067-32-7,
    Sodium hyaluronate
    RL: MOA (Modifier or additive use); TEM (Technical or engineered material
    use); THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (porous .beta.-tricalcium phosphate granules for bone implantation)
     9004-61-9 HCAPLUS
RN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
    9007-28-7 HCAPLUS
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
         1
    CRN
         9007-27-6
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
         2
    CRN
         7664-93-9
    CMF H2 O4 S
     OH
     9067-32-7 HCAPLUS
RN
```

Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

CN

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*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 3 OF 15 HCAPLUS COPYRIGHT 2003 ACS
L75
     2002:533183 HCAPLUS
ΑN
DN
     137:83679
     Pharmaceutical compositions containing viscoelastic substances
TI
     and chemical agents
ΙN
     Tanaka, Koichiro
     Kobayakawa, Shinichiro, Japan; Shinoda, Takuya
PΑ
SO
     Jpn. Kokai Tokkyo Koho, 9 pp.
     CODEN: JKXXAF
DT
     Patent
LA
     Japanese
IC
     ICM A61K045-08
         A61K009-08; A61K047-36; A61K047-38; A61P027-02; A61P029-00;
          A61P031-00
     63-6 (Pharmaceuticals)
FAN.CNT 1
                                    date! APPLICATION NO.
                                                            DATE
                      KIND DATE
     PATENT NO.
                                                                        introcular!
                                           _____
                                                           _____
                           20020716
                                           JP 2001-120001
                                                            20010418
PΙ
     JP 2002201143
                      A2
                                                            20020313
                            20021205
                                           US 2002-96715
                      A1
     US 2002183279
PRAI JP 2000-360300
                     A
                            20001023
     JP 2001-120001
                      Α
                            20010418
     The compns., useful for prevention of infection and/or inflammation in
AΒ
     intraocular surgery or injection into joints, contain chem. agents (e.g.,
     antiinflammatory agents and antimicrobial agents) and viscoelastic
     substances. A mixt. of Healon (Na hyaluronate
     ) and 5 .mu.g/mL of Cravit (levofloxacin) significantly inhibited the
     growth of Bacillus subtilis.
     viscoelastic pharmaceutical antiinflammatory antimicrobial
ST
     intraocular surgery; joint injection viscoelastic pharmaceutical
     antimicrobial levofloxacin
ΙT
     Surgery
        (intraocular; pharmaceutical compns. contg. viscoelastic
        substances and antiinflammatory and/or antimicrobial agents for)
TT
     Anti-inflammatory agents
     Antibacterial agents
     Antibiotics
     Antimicrobial agents
       Viscoelastic materials
        (pharmaceutical compns. contg. viscoelastic substances and
        antiinflammatory and/or antimicrobial agents)
IT
     Cataract
       Eye
        (surgery; pharmaceutical compns. contg. viscoelastic
        substances and antiinflammatory and/or antimicrobial agents for)
IT
     70458-96-7, Norfloxacin
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (Noflo; pharmaceutical compns. contg. viscoelastic substances
        and antiinflammatory and/or antimicrobial agents)
IT
     9067-32-7, Healon
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (Opegan; pharmaceutical compns. contg. viscoelastic
        substances and antiinflammatory and/or antimicrobial agents)
     859-18-7, Lincomycin hydrochloride 82419-36-1, Ofloxacin
ΙT
                                                                 100986-85-4,
     RL: BSU (Biological study, unclassified); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (pharmaceutical compns. contg. viscoelastic substances and
        antiinflammatory and/or antimicrobial agents)
```

```
9004-65-3,
ΙT
    9004-61-9, Hyaluronic acid
    Hydroxypropyl methyl cellulose 9007-28-7, Chondroitin
    sulfate 9082-07-9, Sodium chondroitin
     sulfate 123352-36-3, Viscoat
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical compns. contg. viscoelastic substances and
        antiinflammatory and/or antimicrobial agents)
     9067-32-7, Healon
ΙT
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (Opegan; pharmaceutical compns. contg. viscoelastic
        substances and antiinflammatory and/or antimicrobial agents)
RN
     9067-32-7 HCAPLUS
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9004-61-9, Hyaluronic acid 9007-28-7
     , Chondroitin sulfate 9082-07-9, Sodium
    chondroitin sulfate 123352-36-3, Viscoat
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (pharmaceutical compns. contg. viscoelastic substances and
        antiinflammatory and/or antimicrobial agents)
RN
     9004-61-9 HCAPLUS
CN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9007-28-7 HCAPLUS
RN
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
    CM
         9007-27-6
    CRN
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          2
    CM
    CRN
         7664-93-9
    CMF H2 O4 S
   0
HO-S
     OH
   0
RN
    9082-07-9 HCAPLUS
    Chondroitin, hydrogen sulfate, sodium salt (9CI) (CA INDEX NAME)
CN
    CM
          1
    CRN
         9007-27-6
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          2
    CM
    CRN 7664-93-9
```

CMF H2 O4 S

RN 123352-36-3 HCAPLUS

CN Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid sodium salt (9CI) (CA INDEX NAME)

CM 1

CRN 9067-32-7

CMF Unspecified

CCI PMS, MAN

# \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 9082-07-9

CMF H2 O4 S . x Na . x Unspecified

CM 3

CRN 9007-27-6

CMF Unspecified

CCI PMS, MAN

#### \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4

CRN 7664-93-9

CMF H2 O4 S

L75 ANSWER 4 OF 15 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:429542 HCAPLUS

DN 137:11003

TI Chondroprotective/restorative compositions containing hyaluronic

IN Pierce, Scott W.

PA USA

SO U.S. Pat. Appl. Publ., 14 pp. CODEN: USXXCO

DT Patent

LA English

IC ICM A61K031-715 ICS A61K031-70

NCL 514054000

```
63-6 (Pharmaceuticals)
CC
    Section cross-reference(s): 1, 17
FAN.CNT 1
    PATENT NO.
                   KIND DATE
                                          APPLICATION NO. DATE
                     ----
    -----
                                          -----
    US 2002068718
                     A1
                           20020606
                                          US 2001-967977
                                                           20011002 4
PI
PRAI US 2000-237838P P
                           20001003
    An oral compn. based on hyaluronic acid or its salts
    and optionally a therapeutic drug is provided for treating or preventing
    osteoarthritis, joint effusion, joint inflammation and pain, synovitis,
    lameness, post-operative arthroscopic surgery, deterioration of proper
    joint function including joint mobility, the redn. or inhibition of
    metabolic activity of chondrocytes, the activity of enzymes that degrade
    cartilage, and the redn. or inhibition of the prodn. of hyaluronic
    acid in a mammal. Addnl., compns. contg. hyaluronic
    acid, chondroitin sulfate and glucosamine
    sulfate in a paste formulation are also described which can be
    administered on their own or can be used as a feed additive for cats and
    dogs. For example, a compn. contained (by wt.) glucosamine
    sulfate 36%, chondroitin sulfate 4%,
    sodium hyaluronate 0.144%, manganese sulfate
    0.144%, ibuprofen 200 mg, powd. sugar 20%, glycerin 0.7%, xanthan gum
    0.2%, sodium benzoate 0.7%, citric acid 0.2%, molasses 23.5%, and water
    14.4%.
ST
    oral hyaluronic acid chondrocyte cartilage joint
    disorder; antiarthritic oral hyaluronic acid
    chondrocyte cartilage
TT
    Balsams
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (Peru; chondroprotective/restorative compns. contq. hyaluronic
       acid for treatment of joint disorders)
TT
    Natural products, pharmaceutical
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (aloe; chondroprotective/restorative compns. contg. hyaluronic
       acid for treatment of joint disorders)
    Caseins, biological studies
IT
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (calcium complexes; chondroprotective/restorative compns. contg.
       hyaluronic acid for treatment of joint disorders)
IT
    Drug delivery systems
        (capsules; chondroprotective/restorative compns. contg.
       hyaluronic acid for treatment of joint disorders)
IT
    Natural products, pharmaceutical
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (cascara sagrada; chondroprotective/restorative compns. contg.
       hyaluronic acid for treatment of joint disorders)
ΙT
    Analgesics
    Anti-inflammatory agents
    Antiarthritics
    Cat (Felis catus)
    Dog (Canis familiaris)
    Feed additives
    Horse (Equus caballus)
    Mammalia
    Molasses
    Nutrients
    Witch hazel
        (chondroprotective/restorative compns. contg. hyaluronic
       acid for treatment of joint disorders)
IT
    Amino acids, biological studies
    Castor oil
    Cocoa butter
```

Cod liver oil

Hydrocarbon oils Kaolin, biological studies Lanolin Lecithins Mineral elements, biological studies Sulfonamides Vitamins RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) IT (degrdn. of; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) IT Joint, anatomical (disease, effusion; chondroprotective/restorative compns. contq. hyaluronic acid for treatment of joint disorders) IΤ Leq (disease, lameness; chondroprotective/restorative compns. contq. hyaluronic acid for treatment of joint disorders) ΙT Drug delivery systems (gels; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) ΙT Natural products, pharmaceutical RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (ipecac; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) ITDrug delivery systems (oral; chondroprotective/restorative compns. contq. hyaluronic acid for treatment of joint disorders) IT Drug delivery systems (pastes; chondroprotective/restorative compns. contq. hyaluronic acid for treatment of joint disorders) IT Essential oils RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (peppermint; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) Fatty acids, biological studies ΙT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (polyunsatd., n-3; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) IΤ Surgery (post-operative arthroscopic surgery; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) ΙT Chondrocyte (redn. or inhibition of metabolic activity of; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) Fats and Glyceridic oils, biological studies IT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (sesame; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) Fats and Glyceridic oils, biological studies ΙT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (shark-liver oil; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) IT Synovial membrane (synovitis; chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) 9004-61-9, Hyaluronic acid 9007-28-7 TT , Chondroitin sulfate 9067-32-7, Sodium hyaluronate 29031-19-4, Glucosamine sulfate

RL: FFD (Food or feed use); PAC (Pharmacological activity); THU

(Therapeutic use); BIOL (Biological study); USES (Uses) (chondroprotective/restorative compns. contq. hyaluronic acid for treatment of joint disorders) 50-03-3, Hydrocortisone acetate 50-06-6, Phenobarbital, IT 50-02-2 biological studies 50-13-5, Meperidine hydrochloride 50-21-5, Lactic acid, biological studies 50-23-7, Hydrocortisone 50-24-8, Prednisolone 50-33-9, Phenylbutazone, biological studies 50-78-2, Acetylsalicylic 50-78-2D, Acetylsalicylic acid, buffered 50-81-7, L-Ascorbic acid, biological studies 51-42-3, Epinephrine bitartrate 51-98-9, Norethindrone acetate 52-28-8, Codeine phosphate 53-03-2, Prednisone 53-86-1, Indomethacin 54-11-5, Nicotine 54-31-9, Furosemide 55-63-0 55-63-0, Nitroglycerin 56-75-7, Chloramphenicol 56-81-5, Glycerin, biological studies 57-11-4, Stearic acid, biological studies 57-27-2, Morphine, biological studies 57-33-0, Pentobarbital sodium 57-41-0, Phenytoin 57-55-6, Propylene glycol, biological studies 57-63-6, Ethinyl estradiol 58-08-2, Caffeine, biological studies 58-55-9, Theophylline, biological studies 58-85-5, Biotin 58-93-5, Hydrochlorothiazide 59-30-3, Folic acid, biological studies 59-43-8, Thiamine, biological studies 59-67-6, Niacin, biological studies 61-33-6, biological studies 61-68-7, Mefenamic acid 61-76-7, Phenylephrine hydrochloride 62-49-7, Choline 64-17-5, Ethanol, biological studies 64-19-7, Acetic acid, 64-75-5, Tetracycline hydrochloride 65-23-6, biological studies 65-85-0, Benzoic acid, biological studies 67-63-0, Pyridoxine Isopropanol, biological studies 67-68-5, Dimethyl sulfoxide, biological 67-71-0, Methylsulfonylmethane 68-04-2, Sodium citrate studies 68-19-9, Cyanocobalamin 68-22-4, Norethindrone 69-53-4, Ampicillin 69-72-7, Salicylic acid, biological studies 71-58-9, Medroxyprogesterone acetate 73-78-9, Lidocaine hydrochloride 76-22-2, Camphor 76-49-3, Bornyl acetate 76-57-3, Codeine 77-09-8, Phenolphthalein 77-41-8, 77-92-9, Citric acid, biological studies 78-11-5, Methsuximide Pentaerythritol tetranitrate 79-83-4 83-88-5, Riboflavin, biological 85-79-0, Dibucaine 87-67-2, Choline bitartrate, biological studies 88-04-0, Chloroxylenol 89-78-1, 87-89-8, myo-Inositol studies 90-64-2 93-14-1, Guaifenesin 93-60-7, Methyl nicotinate 94-09-7, Benzocaine 94-36-0, Benzoyl peroxide, biological studies 97-59-6, Allantoin 98-92-0, Niacinamide 100-97-0, Methenamine, biological studies 103-90-2, Acetaminophen 104-46-1, Anethole 108-46-3, Resorcinol, biological studies 108-95-2, Phenol, biological studies 112-38-9, Undecylenic acid 113-92-8, Chlorpheniramine maleate 114-07-8, Erythromycin 115-67-3, Paramethadione 117-10-2, Danthron 119-36-8, Methyl salicylate 119-61-9D, Benzophenone, derivs. 123-03-5, Cetylpyridinium chloride 124-94-7, Triamcinolone 125-69-9, Dextromethorphan hydrobromide 126-07-8, Griseofulvin 128-49-4, Docusate calcium 131-53-3, Dioxybenzone 131-57-7, Oxybenzone 132-20-7, Pheniramine maleate 134-31-6, 8-Hydroxyquinoline sulfate 136-77-6, Hexylresorcinol 137-58-6, Lidocaine 139-12-8, Aluminum acetate 140-65-8, Pramoxine 141-01-5, Ferrous fumarate 143-71-5, Hydrocodone bitartrate 144-55-8, Sodium bicarbonate, biological studies 147-24-0, Diphenhydramine hydrochloride 150-13-0, p-Aminobenzoic acid 152-11-4, Verapamil hydrochloride 152-43-2, Quinestrol 154 - 41 - 6, Phenylpropanolamine hydrochloride 156-51-4, Phenelzine sulfate 299-29-6, Ferrous gluconate 299-42-3, Ephedrine 302-79-4, Tretinoin 303-25-3, Cyclizine hydrochloride 318-98-9, Propranolol hydrochloride 321-64-2, Tacrine 345-78-8, Pseudoephedrine hydrochloride 395-28-8 439-14-5, Diazepam 443-48-1, Metronidazole 469-62-5, Propoxyphene 470-82-6, Eucalyptol 471-34-1, Calcium carbonate, biological studies 532-32-1, Sodium benzoate 546-93-0, Magnesium 532-03-6, Methocarbamol 550-70-9, Triprolidine hydrochloride 557-04-0, Magnesium carbonate e 557-08-4, Zinc undecylenate 562-10-7 577-11-7, Docust 603-50-9, Bisacodyl 614-39-1, Procainamide hydrochloride 577-11-7, Docusate stearate 637-07-0, Clofibrate 637-58-1, Pramoxine hydrochloride Meclofenamic acid 723-46-6, Sulfamethoxazole 980-71-2,

1218-35-5, Xylometazoline hydrochloride Bromopheniramine maleate 1305-62-0, Calcium hydroxide, biological studies 1309-42-8, Magnesium hydroxide 1321-11-5, Aminobenzoic acid 1327-41-9, Aluminum 1400-61-9, Nystatin 1403-66-3, Gentamicin 1404-90-6, chlorohydrate 1405-10-3, Neomycin sulfate 1405-20-5, Polymyxin B sulfate Vancomycin 1405-41-0, Gentamycin sulfate 1405-87-4, Bacitracin 1406-16-2, Vitamin 1639-60-7, Propoxyphene hydrochloride 1406-18-4, Vitamin E 1684-40-8, Tacrine hydrochloride 2391-03-9, Dexbrompheniramine maleate 2955-38-6, Prazepam 3380-34-5, Triclosan 2398-96-1, Tolnaftate 4205-91-8, Clonidine hydrochloride 4499-40-5, 4205-90-7, Clonidine Oxtriphylline, biological studies 5466-77-3, Octyl methoxycinnamate 5534-09-8, Beclomethasone dipropionate 5874-97-5, Metaproterenol sulfate 6385-02-0, Meclofenamate sodium 6740-88-1, Ketamine 7054-25-3Quinidine gluconate 7280-37-7, Estropipate 7439-89-6, Iron, biological 7440-50-8, Copper, 7439-96-5, Manganese, biological studies biological studies 7440-66-6, Zinc, biological studies 7440-70-2, Calcium, biological studies 7447-40-7, Potassium chloride, biological 7460-12-0, Pseudoephedrine sulfate 7491-09-0, Docusate 7553-56-2, Iodine, biological studies 7631-86-9, Silicon potassium dioxide, biological studies 7647-14-5, Sodium chloride (NaCl), biological studies 7681-49-4, Sodium fluoride, biological studies 7704-34-9, Sulfur, biological studies 7720-78-7, Ferrous sulfate 7723-14-0, Phosphorus, biological studies 7733-02-0, Zinc sulfate 7757-79-1, Potassium nitrate, biological studies 7785-87-7, Manganese 8011-96-9, Calamine 8025-63-6 8050-81-5, Simethicone 8065-29-0, Liotrix 9004-10-8, Insulin, biological studies 9004-32-4, Sodium carboxymethyl cellulose 9004-67-5, Methyl cellulose 9005-25-8, 9006-65-9, Dimethicone 9036-19-5, Octoxynol Starch, biological studies 10163-15-2, Sodium monofluorophosphate 11041-12-6, Cholestyramine resin 11096-26-7, Erythropoietin 11099-07-3, Glyceryl stearate 11103-57-4, Vitamin A 11111-12-9D, Cephalosporin, derivs. 11138-66-2, Xanthan gum 12001-79-5, Vitamin K 14362-31-3, Chlorcyclizine 12001-76-2, Vitamin B 14455-29-9, Aluminum carbonate 14663-23-1, Dantrium hvdrochloride 14987-04-3, 14698-29-4, Oxolinic acid 14838-15-4, Phenylpropanolamine Magnesium trisilicate 15307-79-6, Diclofenac sodium 15686-71-2, Cephalexin 15687-27-1, Ibuprofen 17140-78-2, Propoxyphene napsylate 18472-51-0, Chlorhexidine gluconate 18559-94-9, Albuterol 18917-89-0, Magnesium salicylate 20830-75-5, Digoxin 21245-02-3, Padimate O 21645-51-2, Aluminum hydroxide, biological studies 21829-25-4, Nifedipine 22204-53-1, Naproxen 22832-87-7, Miconazole nitrate 22839-47-0, Aspartame 24390-14-5, Doxycycline hyclate 25441-16-1 25812-30-0, Gemfibrozil 26027-38-3, Nonoxynol-9 26159-34-2, Naproxen 26171-23-3, Tolmetin 26787-78-0, Amoxicillin 26921-17-5, Timolol maleate 28911-01-5, Triazolam 28981-97-7, Alprozolam 29094-61-9, Glipizide 29122-68-7, Atenolol 29984-33-6, Vidarabine phosphate 34552-84-6, Isoxicam 34580-13-7, Ketotifen RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (chondroprotective/restorative compns. contg. hyaluronic acid for treatment of joint disorders) 36653-82-4, Cetyl alcohol 36322-90-4, Piroxicam 36505-84-7, Buspirone 37148-27-9, Clenbuterol 38304-91-5, Minoxidil 42399-41-7, Diltiazem 42461-84-7, Flunixin Meglumine 50370-12-2, Cefadroxil 50679-08-8, 51022-70-9, Albuterol sulfate 51264-14-3, Amsacrine Terfenadine 52618-67-4, Tioperidone 53910-25-1, 52128-35-5, Trimetrexate 56296-78-7, Fluoxetine hydrochloride 53994-73-3, Cefaclor Pentostatin 56392-17-7, Metoprolol tartrate 59729-33-8, Citalopram 60142-96-3, Gabapentin 62571-86-2, Captopril 66357-35-5, Ranitidine 68252-19-7, 69198-10-3, Metronidazole 68497-62-1, Pramiracetam Pirmenol 70059-30-2, Cimetidine hydrochloride 72332-33-3, hydrochloride 74011-58-8, Enoxacin 75330-75-5, 73590-58-6, Omeprazole Procaterol 76547-98-3, Lisinopril 80841-47-0, Lovastatin 75847-73-3, Enalapril 88637-37-0, Diphenhydramine citrate 85441-61-8, Quinapril Amsalog 89197-32-0, Efaroxan 93107-08-5, Ciprofloxacin hydrochloride

ΙT

93738-40-0, Ralitoline

93390-81-9, Fosphenytoin

96328-17-5,

```
2'-Chloropentostatin
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (chondroprotective/restorative compns. contq. hyaluronic
        acid for treatment of joint disorders)
ΙT
     9004-34-6, Cellulose, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (microcryst.; chondroprotective/restorative compns. contg.
        hyaluronic acid for treatment of joint disorders)
ΙT
     9004-61-9, Hyaluronic acid 9007-28-7
     , Chondroitin sulfate 9067-32-7,
     Sodium hyaluronate
     RL: FFD (Food or feed use); PAC (Pharmacological activity); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (chondroprotective/restorative compns. contg. hyaluronic
        acid for treatment of joint disorders)
RN
     9004-61-9 HCAPLUS
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9007-28-7 HCAPLUS
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
    CM
          1
    CRN
         9007-27-6
    CMF
          Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          2
    CM
         7664-93-9
    CRN
    CMF
         H2 O4 S
  - s- oh
   0
RN
     9067-32-7 HCAPLUS
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 5 OF 15 HCAPLUS COPYRIGHT 2003 ACS
L75
ΑN
     2000:869032 HCAPLUS
DN
     135:55996
    Healon5 versus Viscoat during cataract surgery: Intraocular pressure,
TΙ
     laser flare and corneal changes
     Schwenn, Oliver; Dick, H. Burkhard; Krummenauer, Frank; Christmann,
ΑU
     Stefan; Vogel, Annette; Pfeiffer, Norbert
     Department of Ophthalmology, Johannes Gutenberg University, Mainz, 55131,
CS
     Graefe's Archive for Clinical and Experimental Ophthalmology (2000),
SO
     238(10), 861-867
     CODEN: GACODL; ISSN: 0721-832X
PB
     Springer-Verlag
DT
     Journal
```

- LA English
- CC 1-12 (Pharmacology)
- The use of a viscoelastic substance facilitates cataract surgery. AB Healon 5 is a new viscoelastic product with special rheol. properties. We evaluated the postoperative effect of Viscoat and Healon5 on intraocular pressure (IOP), central corneal thickness (CCT), endothelial cell counts and laser flare. Forty-eight eyes of 48 patients undergoing routine phacoemulsification followed by foldable IOL implantation were enrolled. Either Healon5 or Viscoat was used according to a block-randomization scheme. The aspiration technique was standardized. IOP, CCT, endothelial cell counts and laser flare were compared pre-and postoperatively. Statistical anal. was performed using the two-sample Wilcoxon test. Data description was based on median and quartiles, while graphic description was performed by non-parametric box plots. Viscoat demonstrated a statistically significant higher IOP than Healon5 at 4 and 8 h postoperatively (P<0.01 and <0.05, resp.). Further, the laser flare values were statistically significantly higher for the Viscoat than for the Healon5 group 8 h postoperatively (P<0.05). Endothelial cell loss did not differ significantly between the two groups (relative change in endothelial cell d. after 3 mo: -4.3% for the Healon5 group and -6.2% for Viscoat group). There was neither a statistically nor a clin. significant difference in endothelial cell loss after the use of Healon5 or Viscoat in routine cataract surgery. However, the IOP in the early postoperative period was higher in the Viscoat group than in the Healon5 group.
- ST Healon5 Viscoat cataract surgery laser flare cornea; intraocular pressure antiglaucoma Healon cataract surgery
- IT Antiglaucoma agents

(Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT Surgery

(cataract; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT Eye

(cornea; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT Laser radiation

(flare; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT Cataract

(surgery; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT 9067-32-7, Healon

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(5; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

IT 123352-36-3, Viscoat

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes)

RE.CNT 25 THERE ARE 25 CITED REFERENCES AVAILABLE FOR THIS RECORD RE

- (1) Arshinoff, S; Curr Can Ophthalmic Pract 1989, V7, P1
- (2) Arshinoff, S; J Cataract Refract Surg 1997, V23, P761 MEDLINE
- (3) Arshinoff, S; Ophthalmic Pract 1995, V13, P98
- (4) Arshinoff, S; Ophthalmic Practice 1998, V16, P24
- (5) Auffarth, G; ASCRS Symposium on Cataract, IOL and Refractive Surgery. Book of abstracts 1999, V136
- (6) Dick, H; Ophthalmologe 1999, V96, P193

krishnan - 10 / 082743 (7) Dick, H; Viskoelastika - eine Ubersicht 1998 (8) Glasser, D; Arch Ophthalmol 1986, V104, P1819 HCAPLUS (9) Gross, J; Am J Ophthalmol 1988, V105, P466 MEDLINE (10) Hammer, M; Invest Ophthalmol Vis Sci 1984, V25, P1329 HCAPLUS (11) Hayreh, S; Arch Opthalmol 1980, V98, P1410 MEDLINE (12) Irvine, A; Arch Ophthalmol 1978, V96, P1023 MEDLINE (13) MacRae, S; Am J Ophthalmol 1983, V95, P332 HCAPLUS (14) Miller, D; Viscoelastic materials: basic science and clinical applications 1989, P3 (15) Miyauchi, S; Curr Eye Res 1984, V3, P1063 HCAPLUS (16) Naeser, K; Acta Ophthalmol 1986, V64, P330 MEDLINE (17) Olivius, E; J Am Intraocul Implant Soc 1985, V11, P480 MEDLINE (18) Probst, L; J Cataract Refract Surg 1994, V20, P145 MEDLINE (19) Rainer, G; J Cataract Refract Surg 2000, V26, P271 MEDLINE (20) Ravalico, G; J Cataract Refract Surg 1997, V23, P433 MEDLINE (21) Savage, J; Ophthalmology 1985, V92, P1506 MEDLINE (22) Schubert, H; Exp Eye Res 1984, V39, P137 HCAPLUS (23) Solomon, K; ASCRS Symposium on Cataract, IOL and Refractive Surgery. Book of abstracts 1999, V134 (24) Tetz, M; ASCRS Symposium on Cataract, IOL and Refractive Surgery. Book of abstracts 1999, V7 (25) Vogel, A; ASCRS Symposium on Cataract and Refractive Surgery. Book of abstracts 1999, V134 ΤТ 9067-32-7, Healon RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (5; Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes) 9067-32-7 HCAPLUS RN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME) CN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* 123352-36-3, Viscoat IT RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Healon5 vs. Viscoat during cataract surgery in humans and intraocular pressure, laser flare and corneal changes) RN 123352-36-3 HCAPLUS Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid CN sodium salt (9CI) (CA INDEX NAME) CM 1 CRN 9067-32-7 CMF Unspecified CCI PMS, MAN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* CM 2

CRN 9082-07-9

CMF H2 O4 S . x Na . x Unspecified

CM 3

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4.

CRN 7664-93-9 CMF H2 O4 S

TТ

```
L75 ANSWER 6 OF 15 HCAPLUS COPYRIGHT 2003 ACS
ΑN
     1999:529038 HCAPLUS
DN
     131:139516
     Use of hyaluronidase to reduce viscoelastic-related post-operative
ΤI
     increases in intraocular pressure
     Refojo, Miguel F.; Harooni, Mark; Freilich, Jonathan M.; Abelson, Mark B.
ΙN
PΑ
     The Schepens Eye Research Institute, Inc., USA
SO
     PCT Int. Appl., 25 pp.
     CODEN: PIXXD2
DТ
     Patent
LA
     English
IC
     ICM A61K038-46
     ICS C12N009-26; A01N043-04
CC
     1-12 (Pharmacology)
     Section cross-reference(s): 63
FAN.CNT 1
                                           APPLICATION NO.
     PATENT NO.
                      KIND DATE
                                                            DATE
     ______
                      ____
                            19990819
                                           WO 1999-US3125
                                                            19990212
PΙ
     WO 9940933
                      Α1
        W: AU, CA, JP, US
         RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE
     AU 9926772
                      A1
                            19990830
                                           AU 1999-26772
                                                            19990212
PRAI US 1998-74837P
                     . P
                            19980217
     WO 1999-US3125
                      W
                            19990212
     Small doses, less than 15 IU and preferably less than 10 IU per treated
AB
     eye, of hyaluronidase can safely and effectively be employed to reduce
     postoperative intraocular pressure caused by residual amts. of
     hyaluronan used during anterior segment surgical procedures. The
     hyaluronidase may be administered after surgery, or at 5 IU or less per
     treated eye concomitantly. Hyaluronidase treatment may be combined with
     treatments with other medications.
     hyaluronidase eye surgery intraocular pressure viscoelastic;
    hyaluronan eye surgery intraocular pressure hyaluronidase
TT
     Eye
     Surgery
     Viscoelastic materials
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure)
IT
     Antiglaucoma agents
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure, and use with other agents)
TT
     Drug delivery systems
```

(infusions; hyaluronidase to reduce viscoelastic-related post-operative

(injections; hyaluronidase to reduce viscoelastic-related

post-operative increases in intraocular pressure)

increases in intraocular pressure)

Drug delivery systems

```
Drug delivery systems
ΙT
        (ophthalmic; hyaluronidase to reduce viscoelastic-related
        post-operative increases in intraocular pressure)
IT
     9004-61-9, Hyaluronan 9067-32-7,
     Sodium hyaluronate 123352-36-3, Viscoat
     RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or
     effector, except adverse); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure)
ΙT
     9001-54-1, Hyaluronidase
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure)
     51-84-3, Acetylcholine, biological studies
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure, and use with other agents)
              THERE ARE 4 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE.CNT
RE
(1) Calder; British Journal of Opthalmology 1986, V70(6), P418 MEDLINE
(2) Equi; Journal of Ocular Pharmacology and Therapeutics 1997, V13(4), P289
(3) Lang; Arch Opthalmol 1984, V102(7), P1079 HCAPLUS
(4) Rankova; Documenta Ophthalmologica 1992, V80(6), P381
     9004-61-9, Hyaluronan 9067-32-7,
     Sodium hyaluronate 123352-36-3, Viscoat
     RL: ADV (Adverse effect, including toxicity); BAC (Biological activity or
     effector, except adverse); BSU (Biological study, unclassified); THU
     (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hyaluronidase to reduce viscoelastic-related post-operative increases
        in intraocular pressure)
RN
     9004-61-9 HCAPLUS
CN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9067-32-7 HCAPLUS
CN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
    123352-36-3 HCAPLUS
    Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid
CN
     sodium salt (9CI) (CA INDEX NAME)
    CM
          1
    CRN
          9067-32-7
     CMF
          Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          2
     CM
     CRN 9082-07-9
         H2 O4 S . x Na . x Unspecified
     CMF
```

CM

3

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4

CRN 7664-93-9 CMF H2 O4 S

L75 ANSWER 7 OF 15 HCAPLUS COPYRIGHT 2003 ACS

AN 1998:646491 HCAPLUS

DN 130:32985

TI Efficacy of hyaluronidase in reducing increases in intraocular pressure related to the use of viscoelastic substances

AU Harooni, Mark; Freilich, Jonathan M.; Abelson, Mark; Refojo, Miguel

CS Schepens Eye Research Institute, Harvard Medical School, Boston, MA, USA

SO Archives of Ophthalmology (Chicago) (1998), 116(9), 1218-1221 CODEN: AROPAW; ISSN: 0003-9950

PB American Medical Association

DT Journal

LA English

CC 1-12 (Pharmacology)

To evaluate the efficacy of hyaluronidase in preventing increases in AΒ intraocular pressure related to injections of hyaluronan-contg. viscoelastic substances, 25 white rabbits were divided into 5 groups. groups 1 through 4, 0.15 mL of aq. humor was removed and replaced with 0.10 mL of a viscoelastic substance in both eyes. Addnl., 10 units of hyaluronidase (0.05 mL) was injected in the anterior chamber of the right eye, whereas the left eye was injected with a volumetrically equiv. dose of balanced saline soln. Viscoelastic substances tested were Healon and Healon GV (Pharmacia & Upjohn, Kalamazoo, Mich), Viscoat (Alcon Labs., Fort Worth, Tex), and Ocucoat (Storz Ophthalmics, Clearwater, Fla). In group 5, right eyes were injected with 10 units of hyaluronidase and the left eyes were treated with balanced saline soln. After injections of viscoelastic substance, intraocular pressure rose rapidly, reaching a peak at approx. 46 h after injection and returning to preinjection levels within 24 h. Hyaluronidase significantly decreased intraocular pressure when used with Healon, Healon GV, and Viscoat, but not with Ocucoat. When injected in the absence of viscoelastic, hyaluronidase appeared to decrease intraocular pressure, but this result was not statistically significant. Injections of hyaluronidase into the anterior chamber of rabbits effectively prevent increases in intraocular pressure induced by hyaluronan-contq. viscoelastic substances. This effect may be related to the ability of hyaluronidase to cleave hyaluronan

ST hyaluronidase viscoelastic hyaluronan intraocular pressure

IT Eye

(intraocular pressure; efficacy of hyaluronidase in reducing increases in intraocular pressure related to the use of viscoelastic substances)

IT 9004-61-9, Hyaluronan 9004-65-3 9067-32-7,
 Healon (polysaccharide) 123352-36-3, Viscoat

```
RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (efficacy of hyaluronidase in reducing increases in intraocular
        pressure related to the use of viscoelastic substances)
     9001-54-1, Wydase
TΤ
     RL: BAC (Biological activity or effector, except adverse); BSU (Biological
     study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (efficacy of hyaluronidase in reducing increases in intraocular
        pressure related to the use of viscoelastic substances)
RE.CNT
              THERE ARE 16 CITED REFERENCES AVAILABLE FOR THIS RECORD
RF.
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(2) Barron, B; Am J Ophthalmol 1985, V100, P377 MEDLINE
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(15) Roden, L; Ciba Found Symp 1989, V143, P60 HCAPLUS
(16) Stankiewicz, A; Klin Oczna 1974, V14, P1005 MEDLINE
IT
     9004-61-9, Hyaluronan 9067-32-7,
     Healon (polysaccharide) 123352-36-3, Viscoat
     RL: ADV (Adverse effect, including toxicity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (efficacy of hyaluronidase in reducing increases in intraocular
        pressure related to the use of viscoelastic substances)
RN
     9004-61-9 HCAPLUS
     Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9067-32-7 HCAPLUS
     Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     123352-36-3 HCAPLUS
RN
CN
     Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid
     sodium salt (9CI) (CA INDEX NAME)
     CM
          1
          9067-32-7
     CRN
          Unspecified
     CMF
     CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN
          9082-07-9
         H2 O4 S . x Na . x Unspecified
     CMF
               3
          CM
          CRN
              9007-27-6
```

CMF

Unspecified

CCI PMS, MAN

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM

CRN 7664-93-9 CMF H2 O4 S

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L75 ANSWER 8 OF 15 HCAPLUS COPYRIGHT 2003 ACS
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1997:344803 HCAPLUS ΑN

DN 127:824

TIMethods and means for control of proliferation of remnant epithelial cells following ocular or other surgery

ΙN Gwon, Arlene E.; Hagemeier, Charles J.

Gwon; Arlene E., USA; Hagemeier; Charles J. PΑ

U.S., 9 pp., Cont. of U.S. Ser. No. 463,390, abandoned. SO CODEN: USXXAM

DTPatent

LA English

IC ICM A61K031-70

NCL 514054000

CC 1-12 (Pharmacology)

FAN.CNT 1

KIND DATE APPLICATION NO. DATE PATENT NO. -----\_\_\_\_\_ \_\_\_\_\_\_ US 5627162 19970506 US 1995-374360 19950118 A PΙ PRAI US 1990-463390 19900111

Methods are disclosed for control of undesired cell proliferation of remnant cells following ocular or other surgery, by applying an effective amt. of at least one proteoglycan-type substrate adhesion mol. (SAM) to the site of surgery. SAMs, in particular chondroitin sulfate, hyaluronic acid, and non-toxic,

pharmaceutically acceptable salts thereof, alone or in a compn. form, are typically used to prevent or inhibit growth of lens-related cells in a lens capsule after surgical removal of the lens, or to prevent proliferative vitreoretinopathy following retinal reattachment procedure performed with or without vitrectomy.

surgery epithelial cell proliferation inhibition; proteoglycan substrate ST adhesion mol antiproliferative surgery; eye surgery epithelial cell proliferation inhibition

TΤ Gelatins, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (CM-cellulose delivery vehicle combined with; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT

(lens, capsule; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT

(lens; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

ΙT Proliferation inhibition (proliferation inhibitors; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT Cataract

Epithelium

Intraocular lenses

Surgery

(proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT Fibronectins

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT Eye, disease

Eye, disease

(retina, detachment; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT Eye, disease

(retinopathy, proliferative vitreoretinopathy; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT Eye

Eye

(vitrectomy; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT 9067-32-7, Healon

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Healon; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT 123352-36-3, Viscoat

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Viscoat; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT 9003-01-4, Polyacrylic acid 9004-32-4 9004-65-3, Hydroxypropyl methyl cellulose

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (delivery vehicle; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT 9004-61-9, Hyaluronic acid 9007-28-7

, Chondroitin sulfate

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

IT 9067-32-7, Healon

RL: BAC (Biological activity or effector, except adverse); BSU (Biological study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

(Healon; proteoglycan-type substrate adhesion mol. for remnant epithelial cell proliferation control following ocular or other surgery)

RN 9067-32-7 HCAPLUS

CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

```
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
IT
    123352-36-3, Viscoat
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (Viscoat; proteoglycan-type substrate adhesion mol. for remnant
        epithelial cell proliferation control following ocular or other
        surgery)
RN
    123352-36-3 HCAPLUS
CN
    Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid
    sodium salt (9CI) (CA INDEX NAME)
    CM
          1
    CRN
          9067-32-7
    CMF
          Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
          2
         9082-07-9
    CRN
         {\tt H2~O4~S~.~x~Na~.~x~Unspecified}
    CMF
          CM
               3
          CRN 9007-27-6
          CMF
               Unspecified
          CCI
              PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          CM
               4
          CRN
               7664-93-9
          CMF
              H2 O4 S
      OH
   0
IT
    9004-61-9, Hyaluronic acid 9007-28-7
     , Chondroitin sulfate
    RL: BAC (Biological activity or effector, except adverse); BSU (Biological
    study, unclassified); THU (Therapeutic use); BIOL (Biological study); USES
     (Uses)
        (proteoglycan-type substrate adhesion mol. for remnant epithelial cell
        proliferation control following ocular or other surgery)
RN
     9004-61-9 HCAPLUS
CN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9007-28-7 HCAPLUS
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
```

CM

1

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

L75 ANSWER 9 OF 15 HCAPLUS COPYRIGHT 2003 ACS 1996:483063 HCAPLUS ΑN DN 125:308797 Compatibility and viscosity of sodium hyaluronate and ΤI sodium chondroitin sulfate in the Viscoat formulation ΑU Doshi, Uday; Gan, Owen; Jani, Rajni; Rhone, Erin Formulation Development Department, Alcon Laboratories, Fort CS Worth, TX, 76134, USA Yakuzaigaku (1996), 56(2), 70-77 SO CODEN: YAKUA2; ISSN: 0372-7629 Nippon Yakuzai Gakkai PΒ DT Journal English LA CC 63-5 (Pharmaceuticals) A combination of sodium hyaluronate (HA) and sodium AΒ chondroitin sulfate (CDS) has been detd. to be particularly well suited as a surgical aid during opthalmic surgery.

particularly well suited as a surgical aid during opthalmic surgery. The objective of these studies was to find the amt. of HA required for a specific viscosity for Viscoat viscoelastic soln., and to det. the amt. of CDS that could be combined with HA to yield a homogenous product. First a series of formulations contg. a const. 4% of CDS, and various concns. of HA were measured for viscosity. Data showed that optimum viscosity was obtained for the formulation contg. 3% HA. In the second series, HA was held to a const. 3%, and the CDS concn. varied. These data showed HA and CDS were not homogeneous when the CDS concn. was equal to or greater than 1.67 times the HA concn. Solns. contg. 4% CDS in combination with 3% HA (the concns. found in Viscoat) formed a miscible soln. From these optimization studies, 3% HA and 4% CDS were detd. to be the optimal concns. to obtain the most utilitarian viscosity for Viscoat viscoelastic soln.

ST hyaluronate chondroitin compatibility viscosity Viscoat soln

IT Viscosity

(compatibility and viscosity of **sodium hyaluronate** and sodium **chondroitin sulfate** in Viscoat soln. for opthalmic surgery)

IT Pharmaceutical dosage forms

(solns., viscoelastic; compatibility and viscosity of sodium hyaluronate and sodium chondroitin sulfate in Viscoat soln. for opthalmic surgery)

IT 9067-32-7, Sodium hyaluronate

9082-07-9, Sodium chondroitin sulfate

RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES (Uses)

```
(compatibility and viscosity of sodium hyaluronate
       and sodium chondroitin sulfate in Viscoat soln. for
       opthalmic surgery)
IT
    9067-32-7, Sodium hyaluronate
    9082-07-9, Sodium chondroitin sulfate
    RL: PRP (Properties); THU (Therapeutic use); BIOL (Biological study); USES
        (compatibility and viscosity of sodium hyaluronate
       and sodium chondroitin sulfate in Viscoat soln. for
       opthalmic surgery)
     9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9082-07-9 HCAPLUS
RN
    Chondroitin, hydrogen sulfate, sodium salt (9CI) (CA INDEX NAME)
CN
    CM
    CRN 9007-27-6
    CMF Unspecified
    CCI PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
         2
    CRN 7664-93-9
    CMF H2 O4 S
L75 ANSWER 10 OF 15 HCAPLUS COPYRIGHT 2003 ACS
AN
    1995:546939 HCAPLUS
    122:274105
DN
    Surface-active viscoelastic hyaluronic acid solutions
ΤI
    for ocular use
IN
    Benedetto, Dominick A.
    Escalon Ophthalmics, Inc., USA
PΑ
    PCT Int. Appl., 65 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
     ICM A61K031-70
     ICS A61F013-20; G02C007-02
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
                                         APPLICATION NO. DATE
     PATENT NO.
                     KIND DATE
                     ----
                                         _____
     _____
                     Al 19950316
                                         WO 1994-US10175 19940907
     WO 9507085
PΙ
        W: CA, JP
         RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
PRAI US 1993-116908 19930907
     A modified mucopolysaccharide soln. for use as a biol. active therapeutic
     infusion comprises a pharmaceutical-grade viscoelastic fraction selected
```

from a C3-20 acyl-substituted hyaluronic acid and

mixts. thereof with hyaluronic acid, and hydroxypropylmethylcellulose. These solns. have a surface tension of 40-65 dynes/cm2; the viscoelastic fraction preferably has an av. mol. wt. .gtoreq.50,000. In some embodiments a physiol. buffer fraction is present. The soln. is injected intraocularly to protect internal ocular structures during ocular surgery and to retard aspiration of material from the ocular surgery site. The soln. also can protect internal ocular structures such as corneal endothelium from accidental contact with surgical instruments. Thus, solns. of 2 hyaluronic acid fractions from rooster comb (1 .times. 106 Da at 5 mg/mL and 5 .times. 105 Da at 30 mg/mL) were mixed at a vol. ratio of 2:1. The viscous mixt. easily fractured when suctioned through a 0.3-mm aspiration cannula at a vacuum pressure of 50-200 mm Hg. hyaluronate viscoelastic soln ocular pharmaceutical Surgery (ocular; surface-active viscoelastic hyaluronic acid solns. for ocular use) Viscoelastic materials (surface-active viscoelastic hyaluronic acid solns. for ocular use) Pharmaceutical dosage forms (injections, ophthalmic, surface-active viscoelastic hyaluronic acid solns. for ocular use) 9004-61-9, Hyaluronic acid 9004-61-9D , Hyaluronic acid, esters, fatty 9004-65-3, Hydroxypropylmethylcellulose 9067-32-7 108174-56-7, Amvisc 123352-36-3, Viscoat RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (surface-active viscoelastic hyaluronic acid solns. for ocular use) 9004-61-9, Hyaluronic acid 9004-61-9D , Hyaluronic acid, esters, fatty 9067-32-7 **123352-36-3**, Viscoat RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (surface-active viscoelastic hyaluronic acid solns. for ocular use) 9004-61-9 HCAPLUS Hyaluronic acid (8CI, 9CI) (CA INDEX NAME) \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* 9004-61-9 HCAPLUS Hyaluronic acid (8CI, 9CI) (CA INDEX NAME) \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* 9067-32-7 HCAPLUS Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME) \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* 123352-36-3 HCAPLUS Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid sodium salt (9CI) (CA INDEX NAME) CM 1 CRN 9067-32-7 CMF Unspecified CCI PMS, MAN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* 2 CM

ST

ΙT

TΤ

ΙT

ΤТ

TT

RN

CN

RN

CN

RN

CN

RN

CN

CRN 9082-07-9

```
CMF H2 O4 S . x Na . x Unspecified
```

CM 3

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

## \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4 •

CRN 7664-93-9 CMF H2 O4 S

```
L75 ANSWER 11 OF 15 HCAPLUS COPYRIGHT 2003 ACS
    1994:144233 HCAPLUS
ΑN
DN
    120:144233
TΙ
    Combinations of viscoelastics for use during surgery
IN
    McLaughlin, Richard N.; Lorenzetti, Ole J.
    Alcon Surgical; Inc., USA
PA
    PCT Int. Appl., 18 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
IC
    ICM A61K009-00
CC
    63-7 (Pharmaceuticals)
FAN.CNT 1
    PATENT NO.
                     KIND DATE
                                         APPLICATION NO.
                                                          DATE
                    ----
                                         _____
    _____
                                                         -----
                   · A1
    WO 9325187
                          19931223
                                         WO 1993-US5639
                                                          19930611
PΤ
        W: AU, CA, JP, US
        RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                           19940104
                                       AU 1993-45347
                                                          19930611
    AU 9345347
                     A1
                                         EP 1993-915325
                                                          19930611
    EP 705095
                      Α1
                           19960410
                         19971126
    EP 705095
                     В1
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LI, LU, MC, NL, PT, SE
                                                          19930611
                          19971215
                                       AT 1993-915325
    AT 160504
                     Ē
                      Т3
                                         ES 1993-915325
                                                          19930611
    ES 2110103
                           19980201
PRAI US 1992-897733
                           19920612
                           19930611
    WO 1993-US5639
    Systems for performing surgery, esp. ophthalmic surgery, utilizing
AΒ
    multiple viscoelastic agents with differing physicochem.
    properties are disclosed. The systems enable the skilled surgeon to
    perform certain steps of a surgical procedure with viscoelastic
    agents that are particularly well suited for such steps. A 1st
    viscoelastic agent has greater adherent properties than a 2nd
    viscoelastic agent which has greater cohesive properties than the
    1st agent. Preferably, the 1st agent comprises a combination of
    Na hyaluronate (av. <1000 kDa) .apprx.3 and
    chondroitin sulfate (.apprx.25 kDa) .apprx.4 wt./vol.%
    and the 2nd agent is Na hyaluronate (av. > 2000 kDa)
    at 1.0 wt./vol.%. The viscoelastic agents are useful in
    conducting cataract surgery.
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ST
    viscoelastic material combination eye surgery;
    hyaluronate chondroitin sulfate combination
    cataract surgery
    Collagens, biological studies
IT
    RL: BIOL (Biological study)
        (in system contg. multiple viscoelastic agents, for surgery)
IT
    Eye, disease
        (surgery, system contg. adhesive first viscoelastic agent and
        cohesive second viscoelastic material for)
TT
    Viscoelastic materials
        (system contg. adhesive first and cohesive second, for surgery)
TΤ
    Surgery
        (system contg. adhesive first viscoelastic agent and cohesive
        second viscoelastic material for)
TT
        (system contg. adhesive first viscoelastic agent and cohesive
        second viscoelastic material for surgery of)
TΤ
    Cohesion
        (system contg. viscoelastic agents having properties of
        adhesion and, for surgery)
TΤ
    Adhesion
        (system contg. viscoelastic agents having properties of
        cohesion and, for surgery)
    9003-05-8, Polyacrylamide
                                 9003-39-8, Polyvinylpyrrolidone
                                                                    9004-32-4,
TΤ
                              9004-57-3, Ethylcellulose 9004-65-3,
    Carboxymethylcellulose
    Hydroxypropylmethylcellulose
                                    9004-67-5, Methylcellulose
    9007-28-7, Chondroitin sulfate
    9067-32-7, Sodium hyaluronate
                                     69992-87-6,
    Keratan 153311-76-3
    RL: BIOL (Biological study)
        (in system contg. multiple viscoelastic agents, for surgery)
    9007-28-7, Chondroitin sulfate
IT
    9067-32-7, Sodium hyaluronate
    153311-76-3
    RL: BIOL (Biological study)
        (in system contg. multiple viscoelastic agents, for surgery)
    9007-28-7 HCAPLUS
RN
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
    CM
         1
         9007-27-6
    CRN
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
         2
    CRN 7664-93-9
    CMF H2 O4 S
   0
     - OH
   ·S
RN
    9067-32-7 HCAPLUS
```

Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

CN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

RN 153311-76-3 HCAPLUS

CN Chondroitin, hydrogen sulfate, mixt. with hyaluronic acid sodium salt (9CI) (CA INDEX NAME)

CM 1

CRN 9067-32-7 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 9007-28-7

CMF H2 O4 S . x Unspecified

CM 3

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 4

CRN 7664-93-9 CMF H2 O4 S

L75 ANSWER 12 OF 15 HCAPLUS COPYRIGHT 2003 ACS

AN 1991:457253 HCAPLUS

DN 115:57253

TI Preparation of water-insoluble derivatives of hyaluronic acid as surgical aids and drug delivery systems

IN Burns, James W.; Cox, Steven; Walts, Alan E.

PA Genzyme Corp., USA

SO U.S., 6 pp. CODEN: USXXAM

DT Patent

LA English

IC ICM A61K047-26 ICS C08L001-00

NCL 106162000

CC 63-7 (Pharmaceuticals)

FAN.CNT 5

PATENT NO. KIND DATE APPLICATION NO. DATE \_\_\_\_\_ -----ΡI US 5017229 Α 19910521 US 1990-543163 19900625 WO 9200105 A119920109 WO 1991-US4543 19910625 W: AU, FI, JP, NO RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LU, NL, SE AU 9183924 A1 19920123 AU 1991-83924 19910625

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AU 660282
                       B2
                            19950622
                            19930421
     EP 537292
                       Α1
                                           EP 1991-914691
                                                             19910625
     EP 537292
                       В1
                            19970409
         R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE
                            19931118
                                                             19910625
     JP 05508161
                       Т2
                                           JP 1991-514243
     AT 151294
                            19970415
                       F.
                                           AT 1991-914691
                                                             19910625
                       Т3
                            19970701
                                           ES 1991-914691
                                                             19910625
     ES 2100954
                       В1
                            20010116
                                           US 1992-833973
     US 6174999
                                                             19920211
     NO 9204875
                       Α
                            19921216
                                           NO 1992-4875
                                                             19921216
                       Α
                                           US 1992-997298
     US 5527893
                            19960618
                                                             19921223
     US 5760200
                       A
                            19980602
                                           US 1995-377949
                                                             19950125
     US 6030958
                       Α
                            20000229
                                           US 1997-914320
                                                             19970818
                                           US 1999-376266
     US 6235726
                       B1
                            20010522
                                                             19990818
                                           US 2001-757202
     US 2001039336
                       A1
                            20011108
                                                             20010109
PRAI US 1987-100104
                       Α2
                            19870918
                            19900625
     US 1990-543163
                       Α
                            19910520
     US 1991-703254
                       Α2
     WO 1991-US4543
                            19910625
                       A
     US 1992-833973
                       A3
                            19920211
     US 1994-176334
                       B1
                            19940103
     US 1994-326058
                            19941019
                       A1
     US 1997-914320
                            19970818
                       А3
AΒ
     A biocompatible gel is prepd. by reacting hyaluronic
     acid (I), a polyanionic polysaccharide, and an activating agent
     under conditions sufficient to form the gel. The polysaccharide
     is chosen from the group consisting of CM-cellulose, carboxymethyl
     amylose, chondroitin-6-sulfate, dermatan
     sulfate, heparin and heparin sulfate. The gels
     prevent adhesions or accretions of body tissues during a post-operation or
     healing period. The gels may also include a pharmaceutically
     active substance. Thus, to 100 mL of an ag. soln. (pH 4.7-4.8) contg. I
     0.4% and CNM cellulose 0.19% was added 0.67 g of 1-(3-dimethylaminopropyl)-
     3-ethyl-carbodiimide and the reaction allowed to proceed for 1 h. After
     removal of the ppt. by dialysis against acidified water for 24 h, the
     slurry was cast into flat molds and air-dried for 24 h at room temp.
     membranes were shown to reduce the incidence of postoperative adhesion
     formation in exptl. animal models.
ST
     hyaluronate gel surgical aid; polysaccharide modified
     hyaluronate tissue adhesive; matrix drug hyaluronate
     gel
IT
     Pharmaceutical dosage forms
        (matrix for, polysaccharide-modified hyaluronate gels
        for)
ΙT
     Medical goods
        (polysaccharide-modified hyaluronate gels for, in
        prevention of postoperative tissue adhesion)
ΙT
     Animal tissue
        (postoperative adhesion prevention of, polysaccharide-modified
        hyaluronate gels for)
IT
     Polysaccharides, compounds
     RL: BIOL (Biological study)
        (reaction products, with activated hyaluronates, for surgical
        aids and drug delivery matrixes)
ΙT
     687-64-9DP, L-Lysine methyl ester, reaction products with
     hyaluronate and carbodiimides
                                     1892-57-5DP, reaction products
     with hyaluronate and polysaccharides
                                           2133-40-6DP, L-Proline
     methyl ester hydrochloride, reaction products with hyaluronate
                         2743-40-0DP, L-Leucine ethyl ester hydrochloride,
     and carbodiimides
     reaction products with hyaluronate and carbodiimides
     2748-02-9DP, L-Leucine-tert-butyl ester hydrochloride, reaction products
     with hyaluronate and carbodiimides 6306-52-1DP, L-Valine
```

methyl ester hydrochloride, reaction products with hyaluronate

7517-19-3DP, L-Leucine methyl ester hydrochloride,

and carbodiimides

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reaction products with hyaluronate and carbodiimides
     7524-50-7DP, L-Phenylalanine methyl ester hydrochloride, reaction products
     with hyaluronate and carbodiimides
                                          9004-32-4DP, Sodium
     carboxymethyl cellulose, reaction products with hyaluronate and
     carbodiimides 9004-61-9DP, Hyaluronic acid,
     reaction products with carbodiimides and polysaccharides
                                                                9005-49-6DP,
     Heparin sulfate, reaction products with hyaluronate and
     carbodimides 9067-32-7DP, Sodium hyaluronate
     , reaction products with carbodiimides and polysaccharides
                                                                  10466-61-2DP,
     L-Leucinamide hydrochloride, reaction products with hyaluronate
                        12768-31-9DP, Carboxymethyl amylose, reaction products
     and carbodiimides
                                         18598-74-8DP, L-Isoleucine
     with hyaluronate and carbodiimides
     methyl ester hydrochloride, reaction products with hyaluronate
                         22572-40-3DP, 1-Ethyl-3-(3-
     and carbodiimides
     dimethylaminopropyl)carbodiimide methiodide, reaction products with
     hyaluronate and polysaccharides
                                      22888-59-1DP, L-Arginine methyl
     ester hydrochloride, reaction products with hyaluronate and
                     22888-60-4DP, L-Histidine methylester hydrochloride,
     carbodiimides
     reaction products with hyaluronate and carbodiimides
     24967-94-0DP, reaction products with hyaluronate and
     carbodiimides 25322-46-7DP, reaction products with
     hyaluronate and carbodiimides
     RL: PREP (Preparation)
        (prepn. of, as surgical aids and drug delivery matrixes)
TΤ
     9004-61-9DP, Hyaluronic acid, reaction
     products with carbodiimides and polysaccharides 9067-32-7DP,
     Sodium hyaluronate, reaction products with carbodiimides
     and polysaccharides 25322-46-7DP, reaction products with
     hyaluronate and carbodiimides
     RL: PREP (Preparation)
        (prepn. of, as surgical aids and drug delivery matrixes)
RN
     9004-61-9 HCAPLUS
CN
     Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     9067-32-7 HCAPLUS
RN
     Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     25322-46-7 HCAPLUS
RN
     Chondroitin, 6-(hydrogen sulfate) (9CI) (CA INDEX NAME)
CN
     CM
          1
          9007-27-6
     CRN
     CMF
         Unspecified
     CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN
         7664-93-9
         H2 O4 S
     CMF
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HO-S-OH

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L75 ANSWER 13 OF 15 HCAPLUS COPYRIGHT 2003 ACS
     1989:587541 HCAPLUS
ΑN
     111:187541
DN
     Effect of hyaluronic acid/chondroitin
TΙ
     sulfate on healing of full-thickness tendon lacerations in rabbits
     Meyers, Steven A.; Seaber, Anthony V.; Glisson, Richard R.; Nunley, James
ΑU
     Med. Cent., Duke Univ., Durham, NC, USA
CS
     Journal of Orthopaedic Research (1989), 7(5), 683-9
SO
     CODEN: JOREDR; ISSN: 0736-0266
DT
     Journal
LA
     English
CC
     1-12 (Pharmacology)
AB
     Viscoat, a complex of highly purified high-mol.-wt. hyaluronic
     acid (HA) with chondroitin sulfate (CS), was
     instilled around the rabbit plantar tendon following full-thickness
     laceration and surgical repair. After 3 wk of immobilization, no
     difference in adhesions or tensile strength of the healing tendons existed
     between Viscoat-treated tendons and controls. This contradicts previous
     studies which suggest that hyaluronic acid reduces
     postoperative tendon adhesions.
ST
     tendon laceration healing hyaluronate chondroitin
     sulfate; Viscoat tendon laceration healing
IT
     Tendon
        (disease, injury, healing of, Viscoat effect on)
     9004-61-9D, Hyaluronic acid, compd. with
ΙT
     chondroitin sulfate 9007-28-7D, compd. with
     hyaluronic acid 123352-36-3, Viscoat
     RL: BIOL (Biological study)
        (tendon laceration healing response to)
     9004-61-9D, Hyaluronic acid, compd. with
IT
     chondroitin sulfate 9007-28-7D, compd. with
     hyaluronic acid 123352-36-3, Viscoat
     RL: BIOL (Biological study)
        (tendon laceration healing response to)
     9004-61-9 HCAPLUS
RN
     Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
RN
     9007-28-7 HCAPLUS
     Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
     CM
          1
          9007-27-6
     CRN
     CMF
          Unspecified
     CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN
          7664-93-9
          H2 O4 S
     CMF
   0
```

```
123352-36-3 HCAPLUS
RN
     Chondroitin, hydrogen sulfate, sodium salt, mixt. with hyaluronic acid
CN
     sodium salt (9CI) (CA INDEX NAME)
          1
     CM
     CRN 9067-32-7
     CMF Unspecified
     CCI PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
     CM
          2
     CRN 9082-07-9
         H2 O4 S . x Na . x Unspecified
     CMF
               3
          CM
          CRN
               9007-27-6
          CMF Unspecified
          CCI
               PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          CM
              7664-93-9
          CRN
          CMF H2 O4 S
L75 ANSWER 14 OF 15 HCAPLUS COPYRIGHT 2003 ACS
AN
     1987:188973 HCAPLUS
DN
     106:188973
     Effect of anterior chamber retaining materials on the corneal endothelial
TΤ
     function
ΑU
     Enomoto, Yoshikazu; Kurabuchi, Shinya; Kanagawa, Ryuichi; Matoba, Miho;
     Yamashita, Takayuki
     Dep. Ophthalmol., Wakayama Med. Coll., 640, Japan
CS
SO
     Nippon Ganka Gakkai Zasshi (1986), 90(12), 1474-8
     CODEN: NGZAA6; ISSN: 0029-0203
DT
     Journal
LA
     Japanese
CC
     1-12 (Pharmacology)
AB
     Anterior chamber retaining materials were applied on the corneal
     endothelium of rabbits for 15 min. Hyaluronate [
     9004-61-9], chondroitin sulfate-
     hyaluronate mixt. [108145-77-3], lactated Ringer's
     soln., and air were used as anterior chamber retaining materials. The
```

corneal endothelium was perfused for 2 h with lactated Ringer's soln. in accordance with Dikstein's method. The perfusion conditions were as follows: 1.0~mL/h, 20~mmHg, 37.degree. During irrigation, the corneal endothelium was obsd. every 30 min under specular microscope for corneal thickness measurement and photographing of endothelial cell changes.

```
After irrigation, the cornea was prepd. for SEM. Corneal swelling rates
     (speed) were 26.0 .mu.m/h in hyaluronate, 42.7 .mu.M/h in the
    chondroitin sulfate-hyaluronate, 28.8 .mu.m/h
    in lactated Ringer's soln., and 40.6 .mu.m/h in air.
                                                           Compared with
    lactated Ringer's soln. and air, hyaluronate and the
    chondroitin sulfate-hyaluronate mixt. both had
    very high viscosity. SEM also revealed that the endothelial cells
     appeared to be swollen after application of all anterior chamber retaining
    materials, and only in the chondroitin sulfate-
    hyaluronate mixt., the pit formation was obsd. The results
     demonstrated the advantage of hyaluronate as an anterior chamber
    retaining material.
ST
    cornea endothelium anterior chamber retaining material
IT
    Atmosphere, environmental
        (cornea endothelial function response to)
IT
    Named reagents and solutions
    RL: BIOL (Biological study)
        (Ringer's, lactated, cornea endothelial function response to)
ΙT
        (cornea, function of endothelium of, anterior chamber retaining
       materials effect on)
     Pharmaceutical dosage forms
TT
        (eye solns., anterior chamber retaining materials, cornea endothelial
        function response to)
     9004-61-9 108145-77-3
ΙT
    RL: BIOL (Biological study)
        (cornea endothelial function response to)
ΙT
    9004-61-9 108145-77-3
    RL: BIOL (Biological study)
        (cornea endothelial function response to)
RN
     9004-61-9 HCAPLUS
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    108145-77-3 HCAPLUS
RN
    Chondroitin, hydrogen sulfate, mixt. with hyaluronic acid (9CI)
CN
    NAME)
    CM
    CRN
         9004-61-9
         Unspecified
    CMF
         PMS, MAN
    CCI
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
          2
    CRN
         9007-28-7
     CMF
         H2 O4 S . x Unspecified
          CM
               3
          CRN
               9007-27-6
          CMF
               Unspecified
          CCI
               PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          CM
               4
          CRN 7664-93-9
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CMF H2 O4 S

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о
но-s-он
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ANSWER 15 OF 15 HCAPLUS COPYRIGHT 2003 ACS
ΑN
    1978:486179 HCAPLUS
DN
    89:86179
ΤI
    Hyaluronic acid
    Nakajima, Akimasa
ΙN
PΑ
    Japan
SO
    Jpn. Kokai Tokkyo Koho, 2 pp.
    CODEN: JKXXAF
DT
    Patent
LΑ
    Japanese
IC
    C08B037-00
CC
     6-4 (General Biochemistry)
FAN.CNT 1
                     KIND DATE
    PATENT NO.
                                          APPLICATION NO.
                                                           DATE
     -----
                     ----
                                          _____
                                                           _____
    JP 53037700
                      A2
                           19780406
                                          JP 1976-111983
                                                           19760914
    JP 60009044
                     B4
                            19850307
PRAI JP 1976-111983
                           19760914
    Hyaluronic acid (I) was sepd. by adjusting aq. I
    solns. to pH 1.8-2.8 in the presence of chondroitinsulfuric
    acid (II) to effect the pptn. of I. Thus, 1 kg whale cartilage
    was crushed, 1/8 wt. 50% aq. NaOH was added, the mixt. was stirred 1 h at
    40.degree., made pH 3.5 with HCl, filtered, and the filtrate made pH 2.2
    to ppt. a jelly. The jelly was washed with H2O, dissolved in 0.1M AcONa,
    and a 3-fold vol. of EtOH was added to ppt. 3.1 g pure I Na salt with a
    recovery of 65 g II.
ST
    hyaluronate cartilage purifn
IT
    Cartilage
        (hyaluronic acid of, purifn. and pptn. of)
ΙT
    9007-28-7
    RL: BIOL (Biological study)
        (hyaluronic acid purifn. and pptn. from solns. in
       presence of)
ΙT
    9067-32-7P
    RL: PREP (Preparation)
        (purifn. and pptn. of, of cartilage)
IT
    9007-28-7
    RL: BIOL (Biological study)
        (hyaluronic acid purifn. and pptn. from solns. in
       presence of)
    9007-28-7 HCAPLUS
RN
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
    CM
         1
         9007-27-6
    CRN
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
```

CM 2

CRN 7664-93-9 CMF H2 O4 S

IT 9067-32-7P

RL: PREP (Preparation)

(purifn. and pptn. of, of cartilage)

RN 9067-32-7 HCAPLUS

CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

## => d all hitstr tot 176

L76 ANSWER 1 OF 10 HCAPLUS COPYRIGHT 2003 ACS

AN 2002:487370 HCAPLUS

DN 137:52395

TI Hypertonic ophthalmic irrigating solution adapted for use in LASIK surgery

IN Wang, Pao-Li; Doshi, Uday; Markwardt, Kerry L.; Maddox, Emerson

PA Alcon Universal Ltd., Switz.

SO PCT Int. Appl., 15 pp.

CODEN: PIXXD2

DT Patent

LA English

IC ICM A61K009-08 ICS A61P027-04

CC 63-6 (Pharmaceuticals)

FAN.CNT 1

TAN.CIVI I																			
		PATENT NO.				KI	ND.	DATE			ΑI	PPLI	CATI	ON NC	).	DATE			
	ΡI	WO 2002049612 WO 2002049612			A	2	20020627 20030116			WO 2001-US44534					20011129				
					A.	3													
								KR,											
			RW:	ΑT,	BE,	CH,	CY,	DE,	DK,	ES,	FI,	FR,	GB,	GR,	ΙE,	ΙΤ,	LU,	MC,	ΝL,
				PT,	SE,	TR													
		AU 2002036501			A.	5	2002	0701		AU 2002-36501					20011129			*	
	PRAI	I US 2000-257464P		164P	P		20001220												
		WO 2001-US44534			W		20011129												

AB Sterile, unpreserved, mildly hypertonic ophthalmic solns. and methods for facilitating the closure and sealing of a corneal flap during LASIK surgery are described. The hypertonicity of the solns. causes the corneal flap to contract following the photoablation step of the LASIK procedure, thereby facilitating the fit of the flap upon closure, as well as the adhesion of the flap to adjacent tissues. The soln. may also contain a viscosity-enhancing agent to further promote adhesion and sealing of the corneal flap. For example, an ophthalmic soln. contained (by wt./vol.) chondroitin sulfate 0.1-10%, sodium chloride 0.5-1.0% (to adjust tonicity), potassium chloride 0.075%. calcium chloride 0.048%, magnesium chloride 0.03%, sodium citrate 0.17%, sodium acetate 0.39%, hydrochloric acid/sodium hydroxide as needed for pH 6.5-8.5, and water to 100%.

ST hypertonic ophthalmic soln LASIK surgery

IT Vinyl compounds, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (carboxy-contg., polymers; hypertonic ophthalmic irrigating soln.

. adapted for use in LASIK surgery) ITEye (cornea, flap, contraction of; hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) IT Buffers Electrolytes Laser radiation (hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) Collagens, biological studies ΙT Proteoglycans, biological studies Salts, biological studies RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) ΙT Solutions (hypertonic; hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) Surgery ΙT (laser vision correction; hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) Alcohols, biological studies IT RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (polyhydric; hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) Drug delivery systems IT (solns., ophthalmic; hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) 50-99-7, Dextrose, biological studies 127-09-3, Sodium acetate IT 144-55-8, Sodium bicarbonate, biological studies 1310-73-2, Sodium hydroxide, biological studies 7447-40-7, Potassium chloride, biological 7558-79-4, Dibasic sodium phosphate 7647-01-0, Hydrochloric acid, biological studies 7647-14-5, Sodium chloride, biological studies 7786-30-3, Magnesium chloride, biological studies 9002-89-5, Polyvinyl 9004-34-6D, Cellulose, derivs. 9005-32-7, Alginic acid 9007-28-7, Chondroitin sulfate 9012-76-4, Chitosan 9067-32-7, Sodium hyaluronate 10043-52-4, Calcium chloride, biological studies 11078-30-1, Galactomannan 11138-66-2, Xanthan gum 27025-41-8, Oxidized glutathione 71010-52-1, **Gellan** gum RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) 9007-28-7, Chondroitin sulfate IT 9067-32-7, Sodium hyaluronate RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) (hypertonic ophthalmic irrigating soln. adapted for use in LASIK surgery) 9007-28-7 HCAPLUS RN CN Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME) CMCRN 9007-27-6 Unspecified CMF CCI PMS, MAN \*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\* CM 2 CRN 7664-93-9

CMF H2 O4 S

```
0
HO-S-OH
   0
    9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L76 ANSWER 2 OF 10 HCAPLUS COPYRIGHT 2003 ACS
    2002:487369 HCAPLUS
AΝ
DN
    137:52394
    Ophthalmic lubricating solution adapted for use in LASIK surgery
TΙ
    containing polymers
    Wang, Pao-Li; Jafari, Masoud R.; Markwardt, Kerry L.; Maddox, Emerson
IN
    Alcon Universal Ltd., Switz.; Doshi, Uday
PA
SO
    PCT Int. Appl., 13 pp.
    CODEN: PIXXD2
DΤ
    Patent
LA
    English
    ICM A61K009-08
TC
    ICS A61K047-36
CC
    63-6 (Pharmaceuticals)
FAN. CNT 1
    PATENT NO.
                     KIND DATE
                                          APPLICATION NO. DATE
                     ----
                                          -----
    ______
    WO 2002049611
                     A2 20020627
                                          WO 2001-US44533 20011129
PΙ
        W: AU, BR, CA, JP, KR, US, ZA
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE, TR
    AU 2002032437
                           20020701
                                          AU 2002-32437
                      Α5
                                                           20011129
PRAI US 2000-257304P
                      Ρ
                           20001220 -
                     W
    WO 2001-US44533
                           20011129
    Ocular lubricant solns. adapted to facilitate the formation of a corneal
AB
    flap during LASIK surgery are described. The solns. contain one or more
    viscosity enhancing agents (e.g., chondroitin sulfate
    or cellulose derivs.) in a substantially salt-free, ophthalmically
    acceptable vehicle. For example, a lubricant soln. contained (by wt.)
    chondroitin sulfate 0-10%, glycerin 1-3%, HCl/NaOH as
    needed for pH 6.5-8.5, and water up to 100%.
ST
    polymer ophthalmic soln eye lubricant vision surgery
ΙT
    Viscosity
        (agents for increase of; polymer-based ophthalmic lubricating soln.
       adapted for use in LASIK surgery)
IT
    Vinyl compounds, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (carboxy-contg., polymers; polymer-based ophthalmic lubricating soln.
       adapted for use in LASIK surgery)
ΙT
        (cornea, flap; polymer-based ophthalmic lubricating soln. adapted for
       use in LASIK surgery)
IT
        (laser vision correction; polymer-based ophthalmic lubricating soln.
       adapted for use in LASIK surgery)
IT
    Biopolymers
    Collagens, biological studies
    Proteoglycans, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
```

```
(polymer-based ophthalmic lubricating soln. adapted for use in LASIK
        surgery)
IΤ
     Drug delivery systems
        (solns., ophthalmic; polymer-based ophthalmic lubricating soln. adapted
        for use in LASIK surgery)
     56-81-5, Glycerin, biological studies 9002-89-5, Polyvinyl alcohol
ΙT
     9004-34-6D, Cellulose, derivs. 9004-65-3, Hydroxypropyl methyl cellulose
     9005-32-7, Alginic acid 9007-28-7, Chondroitin
               9012-76-4, Chitosan 9067-32-7, Sodium
                   11078-30-1, Galactomannan 11138-66-2, Xanthan gum
    hyaluronate
     71010-52-1, Gellan gum
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (polymer-based ophthalmic lubricating soln. adapted for use in LASIK
        surgery)
ΙT
     9007-28-7, Chondroitin sulfate
     9067-32-7, Sodium hyaluronate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (polymer-based ophthalmic lubricating soln. adapted for use in LASIK
        surgery)
     9007-28-7 HCAPLUS
RN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
          1
    CM
    CRN
          9007-27-6
     CMF
          Unspecified
    CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
          2
    CM
    CRN
         7664-93-9
    CMF H2 O4 S
   0
     — OН
HO-S-
   0
RN
    9067-32-7 HCAPLUS
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 3 OF 10 HCAPLUS COPYRIGHT 2003 ACS
L76
     2002:487368 HCAPLUS
ΑN
    137:52393
DN
    Ophthalmic irrigating solution adapted for use in lasik surgery
TΤ
     Doshi, Uday; Markwardt, Kerry L.; Wang, Pao-Li; Maddox, Emerson
ΙN
     Alcon Universal Ltd., Switz.
PA
SO
     PCT Int. Appl., 15 pp.
     CODEN: PIXXD2
ĎΤ
     Patent
LA
     English
     ICM A61K009-08
TC
     ICS A61K047-36
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
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APPLICATION NO.

DATE

KIND DATE

PATENT NO.

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WO 2002049610 A2 20020627
                                           WO 2001-US44526 20011129
PΙ
        W: AU, BR, CA, JP, KR, US, ZA
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, TR
                      Α5
                                           AU 2002-30504
                                                            20011129
    AU 2002030504
                            20020701
PRAI US 2000-257571P
                      Ρ
                            20001220
                     W
    WO 2001-US44526
                            20011129
    Compns. and methods for facilitating the formulation and closure of a
AB
    corneal flap during LASIK surgery are described. The compns. and methods
    are based on the use of one or more viscosity-enhancing agents (e.g.,
    chondroitin sulfate) to provide an ophthalmic irrigating
    soln. with improved coating properties and prolonged dwelling time on the
    cornea, thereby providing lubrication for the microkeratome used to form
    the corneal flap, reduce corneal epithelial abrasions, and help to produce
     smooth and consistent cuts with the microkeratome blade.
    eye irrigating soln Lasik surgery chondroitin sulfate
ST
    Vinyl compounds, biological studies
ΙT
    RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (carboxy-contg., polymers; ophthalmic irrigating soln. adapted for use
        in lasik surgery)
     Physiological saline solutions
IΤ
    Viscosity
        (ophthalmic irrigating soln. adapted for use in lasik surgery)
     Proteoglycans, biological studies
IT
    RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (ophthalmic irrigating soln. adapted for use in lasik surgery)
IT
     Drug delivery systems
        (solns., ophthalmic; ophthalmic irrigating soln. adapted for use in
        lasik surgery)
ΙT
        (surgery; ophthalmic irrigating soln. adapted for use in lasik surgery)
     9002-89-5, Polyvinyl alcohol 9004-34-6D, Cellulose, derivs.
TΤ
    Alginic acid 9012-76-4, Chitosan 11078-30-1, Galactomannan
    11138-66-2, Xanthan gum
                              71010-52-1, Gellan gum
    RL: MOA (Modifier or additive use); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (ophthalmic irrigating soln. adapted for use in lasik surgery)
     9007-28-7, Chondroitin sulfate
TT
     9067-32-7, Sodium hyaluronate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ophthalmic irrigating soln. adapted for use in lasik surgery)
     9007-28-7, Chondroitin sulfate
TT
     9067-32-7, Sodium hyaluronate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ophthalmic irrigating soln. adapted for use in lasik surgery)
RN
     9007-28-7 HCAPLUS
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
          1
    CRN
         9007-27-6
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
          2
     CRN 7664-93-9
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CMF H2 O4 S

```
0
HO-S-OH
   0
    9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 4 OF 10 HCAPLUS COPYRIGHT 2003 ACS
L76
ΑN
    2002:332029 HCAPLUS
    136:319437
DN
    Use of hydroxyeicosatetraenoic acid derivatives in intraocular surgery
TI
    Karakelle, Mutlu; Chan, Kwan
IN
PA
    Alcon Universal Ltd., Switz.
SO
    PCT Int. Appl., 28 pp.
    CODEN: PIXXD2
DT
    Patent
    English
LA
    ICM A61K031-202
TC
    ICS A61P041-00; A61L031-04; C07C051-00
CC
    1-12 (Pharmacology)
    Section cross-reference(s): 9, 63
FAN.CNT 1
                                          APPLICATION NO.
                                                            DATE
    PATENT NO.
                      KIND DATE
                                           _____
     ______
                            20020502
                                           WO 2001-US30212
                                                            20010927
PT
    WO 2002034258
                      A1
        W: AU, CA, JP, US
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, TR
    AU 2002014543
                      Α5
                            20020506
                                           AU 2002-14543
                                                            20010927
    US 2002103257
                       A 1
                            20020801
                                           US 2001-965348
                                                            20010927
    US 6462082
                       B2
                            20021008
                       Р
                            20001023
PRAI US 2000-242501P
                      W
                            20010927
    WO 2001-US30212
OS
    MARPAT 136:319437
    The use of HETE derivs. in intraocular surgery (e.g., cataract surgery) is
AB
    disclosed. The HETE derivs. protect and maintain the corneal endothelium.
    hydroxyeicosatetraenoate deriv intraocular surgery cornea endothelium;
ST
    cataract surgery protection cornea endothelium HETE deriv
IT
    Eye
        (cornea, endothelium; hydroxyeicosatetraenoic acid derivs. for
       protection of corneal endothelium during intraocular surgery)
IT
    Drug delivery systems
      Viscoelastic materials
        (hydroxyeicosatetraenoic acid derivs. for protection of corneal
       endothelium during intraocular surgery)
IT
    Surgery
        (intraocular; hydroxyeicosatetraenoic acid derivs. for protection of
        corneal endothelium during intraocular surgery)
TT
    Drug delivery systems
        (ophthalmic; hydroxyeicosatetraenoic acid derivs. for protection of
       corneal endothelium during intraocular surgery)
IT
    Drug delivery systems
        (solns., ophthalmic; hydroxyeicosatetraenoic acid derivs. for
```

protection of corneal endothelium during intraocular surgery)

(solns., surgical irrigating solns.; hydroxyeicosatetraenoic acid

IT

Drug delivery systems

derivs. for protection of corneal endothelium during intraocular

```
surgery)
ΙT
    Cataract
      Eye
        (surgery; hydroxyeicosatetraenoic acid derivs. for protection of
        corneal endothelium during intraocular surgery)
     69845-60-9D, Hydroxyeicosatetraenoic acid, derivs.
IT
    RL: PAC (Pharmacological activity); THU (Therapeutic use); BIOL
     (Biological study); USES (Uses)
        (hydroxyeicosatetraenoic acid derivs. for protection of corneal
        endothelium during intraocular surgery)
    50-99-7, Dextrose, biological studies
                                             64-17-5, Ethanol, biological
ΙT
               144-55-8, Sodium bicarbonate, biological studies
                                                                  1310-73-2,
    Sodium hydroxide, biological studies
                                           7447-40-7, Potassium chloride,
    biological studies
                         7558-79-4, Dibasic sodium phosphate
                                                                7558-80-7,
    Monobasic sodium phosphate
                                  7647-01-0, Hydrochloric acid, biological
               7647-14-5, Sodium chloride, biological studies
                                                                7732-18-5,
    Water, biological studies
                                7786-30-3, Magnesium chloride, biological
     studies 9067-32-7, Hyaluronic acid
     sodium salt 9082-07-9, Sodium
                           10043-52-4, Calcium chloride,
    chondroitin sulfate
                          27025-41-8, Glutathione disulfide
    biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hydroxyeicosatetraenoic acid derivs. for protection of corneal
        endothelium during intraocular surgery)
RE.CNT
              THERE ARE 6 CITED REFERENCES AVAILABLE FOR THIS RECORD
RF.
(1) Alcon Universal Ltd; WO 0134554 A 2001 HCAPLUS
(2) Hecht, G; US 5409904 A 1995 HCAPLUS
(3) Kavoussi, H; US 5103840 A 1992
(4) Meyer, M; DE 19853007 A 2000
(5) Schwartzman, M; US 4906467 A 1990 HCAPLUS
(6) Yanni, J; US 5696166 A 1997 HCAPLUS
    9067-32-7, Hyaluronic acid sodium
TΤ
    salt 9082-07-9, Sodium chondroitin
     sulfate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hydroxyeicosatetraenoic acid derivs. for protection of corneal
        endothelium during intraocular surgery)
     9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9082-07-9 HCAPLUS
RN
    Chondroitin, hydrogen sulfate, sodium salt (9CI) (CA INDEX NAME)
CN
    CM
          1
    CRN
         9007-27-6
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
    CRN
         7664-93-9
     CMF
         H2 O4 S
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L76 ANSWER 5 OF 10 HCAPLUS COPYRIGHT 2003 ACS
AN
    2001:693076 HCAPLUS
    135:231721
DN
    Viscoelastics for use in middle ear surgery
TΤ
IN
    Jafari, Masoud R.; Doshi, Uday
PΑ
    Alcon Universal Ltd., Switz.
SO
    PCT Int. Appl., 21 pp.
    CODEN: PIXXD2
DT
    Patent
LA
    English
    ICM A61K031-00
IC
     63-6 (Pharmaceuticals)
CC
FAN.CNT 1
                     KIND DATE
                                          APPLICATION NO.
                                                           DATE
    PATENT NO.
                     ____
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                                          _____
                                                           _____
                   A2 20010920
A3 20020725
                           20010920
                                          WO 2001-US8064
PI
    WO 2001068079
                                                           20010314
    WO 2001068079
        W: AU, BR, CA, CN, JP, MX, PL, US, ZA
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
             PT, SE, TR
                           20030102
                                          EP 2001-920347
                                                           20010314
    EP 1267894
                      Α2
        R: AT, BE, CH, DE, DK, ES, FR, GB, GR, IT, LI, LU, NL, SE, MC, PT,
             IE, FI, CY, TR
                           20021114
                                          US 2001-857543
                                                           20010606
    US 2002169142
                     A1
PRAI US 2000-189179P
                      P
                           20000314
    WO 2001-US8064
                     W
                           20010314
    Viscoelastic compns. for surgical and non-surgical packing and
AB
    drug delivery, esp., in conjunction with trauma to or disorders of the ear
    are disclosed. Thus, a highly viscous soln. contained HPMC (E4M-K100M)
    2-8, CaCl2 0.048, NaCl 0.525, KCl 0.075, MgCl2 0.030, sodium citrate
    0.170, sodium acetate 0.390 and water to 100%.
    viscoelastic middle ear surgery
ST
IΤ
    Ear
        (middle; viscoelastic materials for middle ear surgery)
TT
    Drug delivery systems
    Surgery
      Viscoelastic materials
    Viscosity
        (viscoelastic materials for middle ear surgery)
    Acrylic polymers, biological studies
TT
    Collagens, biological studies
    Proteoglycans, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic materials for middle ear surgery)
     9003-05-8, Polyacrylamide 9003-39-8, PVP 9003-97-8, Polycarbophil
TT
                                        9004-57-3, Ethyl cellulose
     9004-32-4, Carboxymethyl cellulose
    9004-61-9, Hyaluronic acid 9004-62-0,
    Hydroxyethyl cellulose 9004-65-3, HPMC
                                              9004-67-5, Methyl cellulose
    9007-28-7, Chondroitin sulfate 9012-76-4,
    Chitosan 9067-32-7, Sodium hyaluronate
    106392-12-5, Poloxamer
                             169799-44-4, Keratin (polysaccharide)
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic materials for middle ear surgery)
IT
     9004-61-9, Hyaluronic acid 9007-28-7
```

```
, Chondroitin sulfate 9067-32-7,
    Sodium hyaluronate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic materials for middle ear surgery)
    9004-61-9 HCAPLUS
RN
    Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    9007-28-7 HCAPLUS
RN
CN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
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         9007-27-6
    CRN
    CMF
         Unspecified
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
         2
    CRN 7664-93-9
    CMF H2 O4 S
   0

    OH

   \circ
    9067-32-7 HCAPLUS
RN
CN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L76 ANSWER 6 OF 10 HCAPLUS COPYRIGHT 2003 ACS
ΑN
    2000:420945 HCAPLUS
DN
    133:63951
ΤI
    Viscoelastic compositions containing antioxidants
    Shah, Mandar V.; Doshi, Uday; Markwardt, Kerry L.
IN
    Alcon Laboratories, Inc., USA
PA
    PCT Int. Appl., 34 pp.
SO
    CODEN: PIXXD2
DT
    Patent
LA
    English
    ICM A61K031-00
IC
CC
     63-6 (Pharmaceuticals)
FAN.CNT 1
                     KIND DATE
                                         APPLICATION NO. DATE
    PATENT NO.
     ______
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                                          _____
                                          WO 1999-US29442 19991210←
ΡI
    WO 2000035432
                    A2
                           20000622
    WO 2000035432
                     A3 20001116
        W: AU, BR, CA, JP, US
        RW: AT, BE, CH, CY, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL,
            PT, SE
                     Ρ
                          19981217
PRAI US 1998-112663P
    Compns. and methods for treating mammalian tissues are disclosed.
     compns. have improved stability, and are viscoelastic compns.
     comprising physiol. antioxidants, bifunctional compds. having an
     anti-inflammatory and anti-oxidant moiety covalently linked by an amide or
     ester bond, in a viscoelastic vehicle. The methods are
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particularly useful in the prevention or treatment of inflammatory and
    proliferative events incident to ocular surgery. Thys, a
    viscoelastic compn. contained (S)-6-methoxy-.alpha.-
    methylnaphthaleneacetic acid (R)-2-(3,4-dihydro-6-hydroxy-2,5,7,8-
    tetramethyl-2H-1-benzopyran-2-yl)ethyl ester 0.00146, Cremophor EL 1.0,
     sodium hyaluronate 1.0, dibasic sodium phosphate 0.056,
    monobasic sodium phosphate monohydrate 0.004, NaCl 0.84, sodium ascorbate
     0.025, HCl/NaOH (pH adjustment) and water qs.
    viscoelastic pharmaceutical antioxidant; naphthaleneacetate
ST
    benzopyranethyl ester viscoelastic pharmaceutical
ΙT
    Carboxylic acids, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (benzenecarboxylic; viscoelastic compns. contg. antioxidants)
IT
    Animal tissue
        (disorders; viscoelastic compns. contq. antioxidants)
IT
    Castor oil
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ethoxylated; viscoelastic compns. contq. antioxidants)
IT
    Eye, disease
        (inflammation; viscoelastic compns. contq. antioxidants)
IT
    Anti-inflammatory agents
        (nonsteroidal; viscoelastic compns. contg. antioxidants)
IT
    Antioxidants
    Drug delivery systems
      Eye, disease
      Viscoelasticity
        (viscoelastic compns. contg. antioxidants)
IT
    Collagens, biological studies
    Proteoglycans, biological studies
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic compns. contq. antioxidants)
IT
    50-81-7, Vitamin C, biological studies
                                              50-81-7D, Vitamin C, salts
    53-86-1D, Indomethacin, derivs.
                                       61-68-7D, Mefenamic acid, derivs.
    68-26-8, Vitamin A
                         70-18-8, Reduced glutathione, biological studies
                                  530-78-9D, Flufenamic acid, derivs.
    134-03-2, Sodium ascorbate
     644-62-2D, derivs.
                         1406-18-4, Vitamin E
                                                 4394-00-7D, Niflumic acid,
                                                   7558-79-4, Dibasic sodium
               5104-49-4D, Flurbiprofen, derivs.
                 7558-80-7, Monobasic sodium phosphate
                                                        7647-14-5, Sodium
    phosphate
                                                                9003-39-8, PVP
    chloride, biological studies
                                    9003-05-8, Polyacrylamide
     9004-32-4, Carboxymethyl cellulose sodium salt
                                                      9004-57-3, Ethyl
    cellulose
                 9004-65-3, HPMC
                                   9004-67-5, Methyl cellulose
     9007-28-7, Chondroitin sulfate
     9067-32-7, Sodium hyaluronate
                                     10049-21-5
    13710-19-5D, Tolfenamic acid, derivs.
                                             15307-86-5D, Diclofenac, derivs.
    15687-27-1D, Ibuprofen, derivs.
                                       17969-20-9D, Fenclozic acid, derivs.
    22071-15-4D, Ketoprofen, derivs.
                                        22131-79-9D, Alclofenac, derivs.
                                      22494-42-4D, Diflunisal, derivs.
    22204-53-1D, Naproxen, derivs.
     26171-23-3D, Tolmetin, derivs.
                                      29679-58-1D, Fenoprofen, derivs.
     31793-07-4D, Pirprofen, derivs.
                                       31842-01-0D, Indoprofen, derivs.
     33369-31-2D, Zomepirac, derivs.
                                       34148-01-1D, Clidanac, derivs.
     34645-84-6D, Fenclofenac, derivs.
                                         36330-85-5D, Fenbufen, derivs.
     36616-52-1D, Fenclorac, derivs.
                                       38194-50-2D, Sulindac, derivs.
     40828-46-4D, Suprofen, derivs.
                                      41340-25-4D, Etodolic acid, derivs.
     50270-33-2D, Isofezolac, derivs.
                                        51234-28-7D, Benoxaprofen, derivs.
     51579-82-9D, Amfenac, derivs.
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     53716-49-7D, Carprofen, derivs.
                                       60653-25-0D, Orpanoxin, derivs.
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     Ketorolac, derivs.
                          180344-46-1
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                                                      180344-49-4
                                                                    180344-50-7
     Bromfenac, derivs.
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                   193221-43-1
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                                              276682-92-9
                                                             276682-93-0
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic compns. contg. antioxidants)
TΤ
     9007-28-7, Chondroitin sulfate
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9067-32-7, Sodium hyaluronate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
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    9007-28-7 HCAPLUS
RN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
    CRN
         9007-27-6
         Unspecified
    CMF
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
    CRN
         7664-93-9
    CMF H2 O4 S
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HO-S-OH
   0
    9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
L76 ANSWER 7 OF 10 HCAPLUS COPYRIGHT 2003 ACS
    1998:618373 HCAPLUS
ΑN
DN
    129:265472
    Viscoelastic compositions and methods of use
TI
    Yanni, John M.; Graff, Gustav
IN
PA
    Alcon Laboratories, Inc., USA
    U.S., 11 pp., Cont.-in-part of U.S. 5,607,966.
SO
    CODEN: USXXAM
DT
    Patent
    English
LA
IC
    ICM A61K031-355
    ICS A61K031-34
    514458000
NCL
    63-6 (Pharmaceuticals)
    Section cross-reference(s): 1
FAN.CNT 7
    PATENT NO.
                     KIND DATE
                                          APPLICATION NO.
                                                           DATE
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                      Α
                                          US 1996-768747
                                                           19961217
    US 5811453
                           19980922
PΙ
                                          WO 1997-US22686 19971216 ---
                           19980625
    WO 9826777
                     A1
        W: AU, CA, JP, US
        RW: AT, BE, CH, DE, DK, ES, FI, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
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    AU 728497
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                      A1
                          19991006
                                          EP 1997-951623
                                                          19971216
    EP 946171
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     JP 2001506651
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    US 6242480
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                                                           19990524
PRAI US 1994-368718
                      A2
                           19941223
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US 1994-362718

A2

19941223

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19961217
     US 1996-768747
                       Α
                            19971216
    WO 1997-US22686
                       W
OS
    MARPAT 129:265472
    Compds. and methods for treating ocular tissues are disclosed. The
AB
    methods utilize viscoelastic compns. contg. certain compds.
    having an anti-inflammatory and anti-oxidant moiety covalently linked by
     an amide or ester bond. The compds. are useful in preventing and treating
     inflammatory and proliferative disorders through several mechanisms. Use
     of a surgical soln. contg. 2-(6-hydroxy-2,5,7,8-tetramethyl-3,4-dihydro-2H-
    benzo[1,2-b]pyran-2-yl)ethyl-2-(6-methoxy-2-naphthyl)propionate (I) for
     glaucoma filtration surgery in rabbits ameliorated inflammatory conditions
     resulting from the surgery and decreased bleb failure. A preferred compn.
    contained I 0.000023, Cremophor EL 0.05, Na hyaluronate
     1, Na2HPO4 0.056, NaH2PO4.cntdot.H2O 0.004, NaCl 0.84, HCl q.s., NaOH q.s,
     and water to 100 % wt./vol.
     antiinflammatory antioxidant conjugate viscoelastic ophthalmic
ST
     liq; naproxen deriv hyaluronate ophthalmic soln
IT
    Eye, disease
        (inflammation; viscoelastic compns. contg. compds. with
        anti-inflammatory and antioxidant moieties for treating ocular tissue)
IT
    Viscoelastic materials
        (liq.; viscoelastic compns. contg. compds. with
        anti-inflammatory and antioxidant moieties for treating ocular tissue)
ΙT
     Drug delivery systems
        (solns., ophthalmic; viscoelastic compns. contg. compds. with
        anti-inflammatory and antioxidant moieties for treating ocular tissue)
ΙT
     53-86-1D, Indomethacin;, derivs.
                                      61-68-7D, Mefenamic acid, derivs.
     530-78-9D, Flufenamic acid, derivs.
                                          644-62-2D, derivs. 4394-00-7D,
                              5104-49-4D, Flurbiprofen;, derivs.
                                                                   9004 - 65 - 3,
    Niflumic acid, derivs.
    HPMC 9007-28-7, Chondroitin sulfate
                                     13710-19-5D,
    9067-32-7, Sodium hyaluronate
                               15307-86-5D, Diclofenac., derivs.
    Tolfenamic acid, derivs.
     15687-27-1D, Ibuprofen;, derivs.
                                        22071-15-4D, Ketoprofen;, derivs.
     22131-79-9D, Alclofenac, derivs.
                                        22204-53-1D, Naproxen;, derivs.
     22494-42-4D, Diflunisal;, derivs. 26171-23-3D, Tolmetin, derivs.
                                       30544-47-9D, Etofenamate, derivs.
     29679-58-1D, Fenoprofen;, derivs.
     31793-07-4D, Pirprofen;, derivs.
                                        31842-01-0D, Indoprofen;, derivs.
     33369-31-2D, Zomepirac;, derivs.
                                        34148-01-1D, Clidanac;, derivs.
     34645-84-6D, Fenclofenac;, derivs.
                                          36330-85-5D, Fenbufen;, derivs.
                                        38194-50-2D, Sulindac;, derivs.
     36616-52-1D, Fenclorac;, derivs.
                                       41340-25-4D, Etodolic acid, derivs.
     40828-46-4D, Suprofen;, derivs.
                                         51234-28-7D, Benoxaprofen, derivs.
     50270-33-2D, Isofezolac;, derivs.
                                     52549-17-4D, Pranoprofen;, derivs.
     51579-82-9D, Amfenac;, derivs.
     53716-49-7D, Carprofen;, derivs.
                                        60653-25-0D, Orpanoxin;, derivs.
                                        74103-06-3D, Ketorolac;, derivs.
     68767-14-6D, Loxoprofen;, derivs.
                                         91714-94-2D, Bromfenac;, derivs.
    74711-43-6D, Zaltoprofen, derivs.
                                 180344-49-4
                                               180344-50-7
                                                             193221-42-0
                  180344-47-2
     180344-46-1
     193221-43-1
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (viscoelastic compns. contg. compds. with anti-inflammatory
        and antioxidant moieties for treating ocular tissue)
RE.CNT
              THERE ARE 51 CITED REFERENCES AVAILABLE FOR THIS RECORD
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(2) Anon; DE 3407507 A1 1985 HCAPLUS
(3) Anon; EP 0183869 A1 1986 HCAPLUS
(4) Anon; EP A241043 1987
(5) Anon; EP A279867 1988
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    Res Basel 1990, V5, P1
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     9007-28-7, Chondroitin sulfate
IT
     9067-32-7, Sodium hyaluronate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses) .
        (viscoelastic compns. contg. compds. with anti-inflammatory
        and antioxidant moieties for treating ocular tissue)
RN
     9007-28-7 HCAPLUS
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
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          9007-27-6
    CRN
    CMF
          Unspecified
    CCI
          PMS, MAN
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\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CRN 7664-93-9 CMF H2 O4 S

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RN
     9067-32-7 HCAPLUS
     Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
   ANSWER 8 OF 10 HCAPLUS COPYRIGHT 2003 ACS
L76
     1998:424121 HCAPLUS
ΑN
     129:86034
DN
     Ophthalmic viscoelastic compositions
TI
     Yanni, John M.; Graff, Gustav
ΙN
     Alcon Laboratories, Inc., USA; Yanni, John M.; Graff, Gustav
PΑ
     PCT Int. Appl., 33 pp.
SO
     CODEN: PIXXD2
DT
     Patent
LA
     English
     ICM A61K031-35
IC
     ICS A61K031-24; A61K031-355
     63-6 (Pharmaceuticals)
CC
FAN.CNT 7
     PATENT NO.
                      KIND DATE
                                           APPLICATION NO.
                                                            DATE
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                            19980625
                                           WO 1997-US22686
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                            19980715
                                           AU 1998-55215
                                                            19971216
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     EP 946171
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                                           US 1999-308851
                                                            19990524
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                       Α
PRAI US 1996-768747
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     US 1994-362718
                       A2
                       A2
                            19941223
     US 1994-368718
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W

WO 1997-US22686

OS GI

AΒ

MARPAT 129:86034

19971216

```
antiinflammatory agent linked, either directly or via a linker to a
    benzopyran or benzofuran deriv. in a viscoelastic vehicle.
    Thus, a preferred compn. contains ester I 0.000023, Cremophor EL 0.05,
    hyaluronic acid Na salt 1, Na2HPO4 0.056, NaH2PO4 0.004,
    NaCl 0.84 % w/v in water, pH adjusted with NaOH and HCl.
ST
    antiinflammatory viscoelastic ophthalmic compn
ΙT
    Castor oil
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ethoxylated; ophthalmic viscoelastic compns. contg. a
       non-steroidal antiinflammatory agent)
IT
    Drug delivery systems
        (ophthalmic; ophthalmic viscoelastic compns. contg. a
       non-steroidal antiinflammatory agent)
                           61-68-7, Mefenamic acid
                                                       530-78-9, Flufenamic
    53-86-1, Indomethacin
TΤ
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                      1310-73-2, Sodium hydroxide, biological studies
                                                        7558-79-4, Disodium
    4394-00-7, Niflumic acid
                               5104-49-4, Flurbiprofen
                         7558-80-7, Sodium dihydrogen phosphate
                                                                   7647-01-0,
    hydrogen phosphate
    Hydrochloric acid, biological studies 7647-14-5, Sodium chloride,
                         7732-18-5, Water, biological studies
    biological studies
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    Tolfenamic acid
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    22494-42-4, Diflunisal
                              26171-23-3, Tolmetin
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                                                       31842-01-0, Indoprofen
    30544-47-9, Etofenamate
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                            36616-52-1, Fenclorac
                                                    38194-50-2, Sulindac
    36330-85-5, Fenbufen
                            41340-25-4, Etodolic acid
                                                        50270-33-2, Isofezolac
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    51234-28-7, Benoxaprofen
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                                                      52549-17-4, Pranoprofen
    53716-49-7, Carprofen
                             60653-25-0, Orpanoxin
                                                     68767-14-6, Loxoprofen
                             74711-43-6, Zaltoprofen
    74103-06-3, Ketorolac
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    180344-46-1
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    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ophthalmic viscoelastic compns. contg. a non-steroidal
       antiinflammatory agent)
RE.CNT
             THERE ARE 7 CITED REFERENCES AVAILABLE FOR THIS RECORD
RE
(1) Alcon Laboratories; WO 9620187 A 1996 HCAPLUS
(2) Alcon Laboratories Inc; WO 9710236 A 1997 HCAPLUS
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    9007-28-7, Chondroitin sulfate
IT
    9067-32-7, Sodium hyaluronate
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (ophthalmic viscoelastic compns. contg. a non-steroidal
       antiinflammatory agent)
     9007-28-7 HCAPLUS
RN
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
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         9007-27-6
    CRN
         Unspecified
    CMF
    CCI
         PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
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2

CM

CRN 7664-93-9 CMF H2 O4 S

RN 9067-32-7 HCAPLUS

CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

L76 ANSWER 9 OF 10 HCAPLUS COPYRIGHT 2003 ACS

AN 1995:737582 HCAPLUS

DN 123:123201

TI Combinations of polymers for use in artificial tear compositions

IN Bhagat, Haresh G.

PA Alcon Laboratoires, Inc., USA

SO Eur. Pat. Appl., 7 pp.

CODEN: EPXXDW

DT Patent

LA English

IC ICM A61K009-00 ICS A61K033-10

CC 63-6 (Pharmaceuticals)

FAN. CNT 3

PAN.		-	KIND	DATE	APPLICATI	ON NO.	DATE			
PI			A2 A3	19950719 19951115	EP 1994-6	550039	19941219			
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	CA	R: AT, BE, 2134376	AA	, DK, ES, 19950621	CA 1994-2			ΝL,	Ρ1,	SE
		2134376	C	20011023						
	-	07196474 152348	A2 E	19950801 19970515	JP 1994-2 AT 1994-6		19941129 19941219			
		2102175	Т3	19970716	ES 1994-6		19941219			
		9481604	A1	19950629	AU 1994-8	31604	19941220			
		675371 5460834	B2 A	19970130 19951024	US 1995-3	371043	19950110			
PRAI		1993-170482	A	19931220						
	-	1991-807528 1992-844269	B1 B1	19911213 19920302						
		1992-994051	B2	19921216						
	US	1993-31058	B2 ·	19930312						

- Physiol. tear compns. for the treatment of dry eye syndrome are disclosed which have a high viscosity and contain bicarbonate, a cellulosic polymer and/or a glycosaminoglycan and/or a carboxyvinyl polymer. A claimed compn. with a viscosity 5-10,000 cP comprises: (a) potassium ions 11-25 mmol/L; (b) calcium ions 0.2-0.5 mmol/L; (c) magnesium ions 0.15-0.45 mmol/L; (d) bicarbonate ions 1-36 mmol/L; (e) at least one component from a group consisting a cellulosic polymer, a glycosaminoglycan, and a carboxyvinyl polymer. For example, an eye soln. contained hydroxypropyl Me cellulose 0.5, mannitol q.s., CaCl2.cntdot.2H2O 0.0053, MgCl2.cntdot.6H2O 0.0064, ZnCl2 0.00015, KHCO3 0.1, Carbomer 934P 0.175, and purified water to 100%.
- ST artificial tear dry eye polymer salt
- IT Glycosaminoglycans, biological studies

RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)

```
(salts and polymers for use in artificial tear compns.)
    Tear
IΤ
        (artificial, salts and polymers for use in artificial tear compns.)
    Vinyl compounds, biological studies
TΤ
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (carboxy-contg., polymers, salts and polymers for use in artificial
        tear compns.)
    Eye, disease
IT
        (keratoconjunctivitis sicca, salts and polymers for use in artificial
        tear compns.)
ΙT
     Pharmaceutical dosage forms
        (ophthalmic, salts and polymers for use in artificial tear compns.)
    144-55-8, Sodium bicarbonate, biological studies 298-14-6, Potassium
TT
                   7447-40-7, Potassium chloride, biological studies
    bicarbonate
    7646-85-7, Zinc chloride, biological studies
                                                    7786-30-3, Magnesium
    chloride, biological studies 9004-62-0, Hydroxyethyl cellulose
                                          9004-65-3, Hydroxypropyl methyl
     9004-64-2, Hydroxypropyl cellulose
                 9004-67-5, Methyl cellulose 9007-28-7,
    Chondroitin sulfate 9067-32-7, Sodium
                  10043-52-4, Calcium chloride, biological studies
    hyaluronate
     57916-92-4, Carbomer 934P
                                76050-42-5, Carbomer 940
                                                           91315-32-1,
                    96827-24-6, Carbomer 1342
                                               139637-85-7, Carbomer 980
    Carbomer 910
    RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (salts and polymers for use in artificial tear compns.)
ΙT
     9007-28-7, Chondroitin sulfate
     9067-32-7, Sodium hyaluronate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (salts and polymers for use in artificial tear compns.)
RN
     9007-28-7 HCAPLUS
    Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)
CN
    CM
          1
          9007-27-6
    CRN
    CMF
          Unspecified
    CCI
          PMS, MAN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    CM
          2
         7664-93-9
    CRN
    CMF
         H2 O4 S
     – OН
     9067-32-7 HCAPLUS
RN
    Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)
CN
*** STRUCTURE DIAGRAM IS NOT AVAILABLE ***
    ANSWER 10 OF 10 HCAPLUS COPYRIGHT 2003 ACS
L76
     1995:602342 HCAPLUS
ΑN
     123:17996
DN
     Hyaluronic acid compositions useful as an irrigating
TΙ
     solution in surgery
ΙN
     Hecht, Gerald; Lorenzetti, Ole J.
```

```
PΑ
    Alcon Laboratories, Inc., USA
    U.S., 7 pp. Cont. of U.S. Ser. No. 553,924, abandoned.
SO
    CODEN: USXXAM
DT
     Patent
LA
     English
    ICM A01N043-04
IC
NCL
    514023000
     63-8 (Pharmaceuticals)
CC
FAN. CNT 1
                                           APPLICATION NO.
     PATENT NO.
                      KIND DATE
                                                            DATE
                            _____
     ______
                      ____
                                           ______
    US 5409904
ΡI
                      Α
                            19950425
                                           US 1992-977312
                                                            19921116
    WO 9632929
                            19961024
                                           WO 1995-US4816
                                                            19950419
                      A1
        W: AU, CA, JP, MX
        RW: AT, BE, CH, DE, DK, ES, FR, GB, GR, IE, IT, LU, MC, NL, PT, SE
                                          AU 1995-23895
                                                            19950419
    AU 9523895
                      A1
                            19961107
                                           US 1995-425132
                                                            19950419
    US 5578578
                      Α
                            19961126
PRAI US 1984-671042
                            19841113
    US 1986-899167
                            19860822
    US 1987-95601
                            19870910
    US 1990-553924
                            199.00717 .
    US 1992-977312
                            19921116
    WO 1995-US4816
                            19950419
    Disclosed are solns. useful in surgery comprising a viscous or
AΒ
    viscoelastic substance in an aq. vehicle which is characterized as
    physiol. compatible; also disclosed are methods of using such solns.,
     implanting such viscous or viscoelastic substances, while
    minimizing the traumatic effect of surgery at the cellular level. A soln.
     for use during ocular surgery contained Na hyaluronate
     1, NaCl 1, dried Na phosphate 1, CaCl2 1, MgCl2 1, dextrose 1, glutathione
     0.5, NaHCO3 1, NaOH/HCl q.s. to pH 7.2, and purified water to 100 parts.
    hyaluronate irrigation soln ocular surgery
ST
ΙT
    Collagens, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hyaluronic acid compns. useful as irrigating soln.
        in surgery)
ΙT
        (ocular; hyaluronic acid compns. useful as
        irrigating soln. in surgery)
IT
        (surgery of; hyaluronic acid compns. useful as
        irrigating soln. in surgery)
                                             70-18-8, Glutathione, biological
ΙT
     50-99-7, Dextrose, biological studies
             144-55-8, Sodium bicarbonate, biological studies 7447-40-7,
     studies
     Potassium chloride, biological studies 7632-05-5, Sodium phosphate
     7647-14-5, Sodium chloride, biological studies
                                                      7786-30-3, Magnesium
     chloride, biological studies 9004-61-9, Hyaluronic
            9004-62-0, Hydroxyethyl cellulose
                                               9004-64-2,
     Hydroxypropyl cellulose 9004-65-3, Hydroxypropyl methyl cellulose
     9004-67-5, Methyl cellulose 9007-28-7, Chondroitin
              9062-14-0, Hydroxypropyl ethyl cellulose
     9067-32-7, Sodium hyaluronate 10043-52-4,
     Calcium chloride, biological studies
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hyaluronic acid compns. useful as irrigating soln.
        in surgery)
     9004-61-9, Hyaluronic acid 9007-28-7
ΙT
     , Chondroitin sulfate 9067-32-7,
     Sodium hyaluronate
     RL: THU (Therapeutic use); BIOL (Biological study); USES (Uses)
        (hyaluronic acid compns. useful as irrigating soln.
        in surgery)
RN
     9004-61-9 HCAPLUS
```

CN Hyaluronic acid (8CI, 9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

RN 9007-28-7 HCAPLUS

CN Chondroitin, hydrogen sulfate (9CI) (CA INDEX NAME)

CM 1

CRN 9007-27-6 CMF Unspecified CCI PMS, MAN

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

CM 2

CRN 7664-93-9 CMF H2 O4 S

RN 9067-32-7 HCAPLUS

CN Hyaluronic acid, sodium salt (9CI) (CA INDEX NAME)

\*\*\* STRUCTURE DIAGRAM IS NOT AVAILABLE \*\*\*

=> fil medline

FILE 'MEDLINE' ENTERED AT 17:13:33 ON 15 MAR 2003

FILE LAST UPDATED: 14 MAR 2003 (20030314/UP). FILE COVERS 1958 TO DATE.

On June 9, 2002, MEDLINE was reloaded. See HELP RLOAD for details.

MEDLINE thesauri in the /CN, /CT, and /MN fields incorporate the MeSH 2003 vocabulary. See http://www.nlm.nih.gov/mesh/summ2003.html for a description on changes.

This file contains CAS Registry Numbers for easy and accurate substance identification.

=> d all tot 1124

L124 ANSWER 1 OF 2 MEDLINE

AN 2002265171 MEDLINE

DN 21925045 PubMed ID: 11927686

TI Interactive effect of chondroitin sulphate C and hyaluronan on fluid movement across rabbit synovium.

AU Sabaratnam S; Coleman P J; Badrick E; Mason R M; Levick J R

CS Department of Physiology, St George's Hospital Medical School, London SW17 ORE, UK.

SO JOURNAL OF PHYSIOLOGY, (2002 Apr 1) 540 (Pt 1) 271-84. Journal code: 0266262. ISSN: 0022-3751.

CY England: United Kingdom

DT Journal; Article; (JOURNAL ARTICLE)

LA English

FS Priority Journals

ΕM 200209 Entered STN: 20020514 ED Last Updated on STN: 20020924 Entered Medline: 20020923 The polysaccharide hyaluronan (HA) conserves synovial fluid by AB keeping outflow low and almost constant over a wide pressure range ('buffering'), but only at concentrations associated with polymer domain overlap. We therefore tested whether polymer interactions can cause buffering, using HA-chondroitin sulphate C (CSC) mixtures. Also, since it has been found that capillary filtration is insensitive to the Starling force interstitial osmotic pressure in frog mesenteries, this was assessed in synovium. Hyaluronan at non-buffering concentrations (0.50-0.75 mg ml(-1)) and/or 25 mg ml(-1) CSC (osmotic pressure 68 cmH(2)O) was infused into knees of anaesthetised rabbits in vivo. Viscometry and chromatography confirmed that HA interacts with CSC. Pressure (P(j)) versus trans-synovial flow (;Q(s)) relations were measured.; Q(s) was outwards for HA alone (1.2 +/- 0.9 microl min(-1) at 3 cmH(2)0, mean +/- S.E.M.; n = 6). CSC diffused into synovium and changed; Q(s) to filtration at low P(j) (-4.1 microl min(-1), 3 cmH(2)0, n = 5, P < 0.02, t test). Filtration ceased upon circulatory arrest (n = 3). At higher P(j), 0.75 mg ml(-1) HA plus CSC buffered; Q(s) to approximately 3 microl min(-1) over a wide range of P(j), with an outflow increase of only 0.04 + - 0.02 microl min(-1) cmH(2)O(-1) (n = 4). With HA or CSC alone, buffering was absent (slopes 0.57 +/- 0.04 microl min(-1) cmH(2)O(-1) (n = 4) and 0.86 +/- 0.05 microl min(-1) cmH(2)O(-1) (n = 5), respectively). Therefore, polymer interactions can cause outflow buffering in joints. Also, interstitial osmotic pressure promoted filtration in fenestrated synovial capillaries, so the results for frog mesentery capillaries cannot be generalised. The difference is attributed to differences in pore ultrastructure. Check Tags: Animal; Support, Non-U.S. Gov't \*Adjuvants, Immunologic: PD, pharmacology Biological Transport: DE, drug effects Capillary Permeability: DE, drug effects \*Chondroitin Sulfates: PD, pharmacology Drug Interactions \*Hyaluronic Acid: PD, pharmacology Knee Joint: ME, metabolism Models, Biological Osmotic Pressure Polymers: ME, metabolism Rabbits \*Synovial Fluid: ME, metabolism Synovial Membrane: BS, blood supply Synovial Membrane: DE, drug effects \*Synovial Membrane: ME, metabolism Viscosity 9004-61-9 (Hyaluronic Acid); 9007-28-7 (Chondroitin RN CN 0 (Adjuvants, Immunologic); 0 (Polymers) L124 ANSWER 2 OF 2 MEDLINE 95271545 ΑN MEDLINE PubMed ID: 7752124 DN 95271545 TΙ Effects of hyaluronic acid on the release of cartilage matrix proteoglycan and fibronectin from the cell matrix layer of chondrocyte cultures: interactions between hyaluronic acid and chondroitin sulfate glycosaminoglycan. ΑU Kato Y; Mukudai Y; Okimura A; Shimazu A; Nakamura S Department of Biochemistry, School of Dentistry, Hiroshima University, CS

JOURNAL OF RHEUMATOLOGY. SUPPLEMENT, (1995 Feb) 43 158-9.

SO

```
Journal code: 7806058. ISSN: 0380-0903.
```

- CY Canada
- DT Journal; Article; (JOURNAL ARTICLE)
- LA English
- FS Priority Journals
- EM 199506
- ED Entered STN: 19950629

Last Updated on STN: 19950629 Entered Medline: 19950622

AB Hyaluronic acid (HA) of large sizes suppressed the release of cartilage matrix proteoglycan, fibronectin, and other macromolecules from the cell matrix layer of chondrocyte cultures, perhaps because HA of large sizes formed a viscous barrier in the matrix by its interactions with other extracellular matrix macromolecules. To test this possibility, we determined the viscosity of solutions containing HA of various sizes in the presence of proteoglycan monomer or chondroitin sulfate glycosaminoglycan (GAG). Not only the monomer but also chondroitin sulfate increased the viscosity of HA solutions, depending on the size of HA. These findings suggest that HA of large sizes increases the viscosity near the surface of articular cartilage by sugar-sugar and by sugar-protein interactions and that the increase of viscosity is involved in the protective action of HA on arthritic cartilage.

CT Check Tags: Animal; Support, Non-U.S. Gov't

Cartilage: CY, cytology
\*Cartilage: DE, drug effects
Cartilage: ME, metabolism
Cells, Cultured

\*Chondroitin Sulfates: PD, pharmacology

Drug Interactions

Fibronectins: DE, drug effects \*Fibronectins: ME, metabolism

\*Hyaluronic Acid: PD, pharmacology Proteoglycans: DE, drug effects \*Proteoglycans: ME, metabolism Rabbits

RN 9004-61-9 (Hyaluronic Acid); 9007-28-7 (Chondroitin Sulfates)

CN 0 (Fibronectins); 0 (Proteoglycans); 0 (chondroitin sulfate glycosaminoglycan)

## => fil embase

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FILE COVERS 1974 TO 13 Mar 2003 (20030313/ED)

EMBASE has been reloaded. Enter HELP RLOAD for details.

This file contains CAS Registry Numbers for easy and accurate substance identification.

## => d all

L153 ANSWER 1 OF 1 EMBASE COPYRIGHT 2003 ELSEVIER SCI. B.V.

AN 94096368 EMBASE

DN 1994096368

- TI Hyaluronic acid. A review of its pharmacology and use as a surgical aid in ophthalmology, and its therapeutic potential in joint disease and wound healing.
- AU Goa K.L.; Benfield P.
- CS Adis International Limited, 41 Centorian Drive, Mairangi Bay, Auckland 10,

```
New Zealand
     Drugs, (1994) 47/3 (536-566).
SO
     ISSN: 0012-6667 CODEN: DRUGAY
CY
     New Zealand
DT
     Journal; General Review
             Ophthalmology
FS
     012
     030
             Pharmacology
             Arthritis and Rheumatism
     031
     033
             Orthopedic Surgery
     037
           · Drug Literature Index
     038
             Adverse Reactions Titles
LA
     English
SL
     English
AΒ
     Hyaluronic acid is a naturally occurring
     polysaccharide with distinct physicochemical properties which underlie its
     application as a viscoelastic tool in ophthalmological surgery. In
     cataract surgery the role of hyaluronic acid in
     facilitating procedures and protecting the corneal endothelium is well
     established. Some benefit was also been gained with the use of
     hyaluronic acid in penetrating keratoplasty
     trabeculectomy retinal reattachment and trauma surgery although its
     efficacy in these indications is less well-defined in the published
     literature. In addition to its lubricating and cushioning properties
     demonstration of some in vitro anti-inflammatory activity and a possible
     disease-modifying effect for hyaluronic acid in
     animals has prompted its investigation as a treatment in osteoarthritis
     and to a much lesser extent in rheumatoid arthritis. Hyaluronic
     acid 20mg as weekly intra-articular injections for 3 to 7 weeks
     improved knee pain and joint motion in patients with osteoarthritis.
     Although this occurred to a greater degree than with placebo in most
     comparisons the effects of hyaluronic acid was similar
     to those of placebo in the largest trial. In the few available comparisons
     with other agents hyaluronic acid appeared equivalent
     to methylprednisolone 40mg (for 3 weeks) and to a single injection of
     triamcinolone 40mg. Hyaluronic acid was distinguished
     from other therapies by providing a sustained effect after treatment
     discontinuation. Together with its very good tolerability profile these
     properties justify further study of hyaluronic acid in
     patients with osteoarthritis. Some limited evidence of improvement in
     patients with rheumatoid arthritis and a possible healing effect of
     hyaluronic acid on tympanic membrane perforations
     represent additional areas of interest for future investigation. In
     summary hyaluronic acid is a well-established adjunct
     to cataract surgery and may prove to be a promising option in the
     treatment of patients with osteoarthritis. Its very good tolerability
     provides further impetus for examination of its potential role in on
     extended scope of arthritic and ophthalmological indications and in wound
     healing.
CT
     Medical Descriptors:
       *arthropathy: DT, drug therapy
     *eye surgery
     *wound healing
       cartilage
     cataract extraction
     clinical trial
     cornea endothelium
     cornea transplantation
     drug binding
     drug blood level
     drug effect
```

drug elimination
drug formulation
drug half life

```
drug structure
eardrum perforation: DT, drug therapy
eye injury: SU, surgery
female
glaucoma: SU, surgery
human
inflammation: SI, side effect
  intraarticular drug administration
intraocular pressure
iritis: SI, side effect
meta analysis
nonhuman
  osteoarthritis: DT, drug therapy
pharmacodynamics
retina surgery
review
  rheumatoid arthritis: DT, drug therapy
  synovial fluid
trabeculectomy
viscoelasticity
Drug Descriptors:
  *hyaluronic acid: AE, adverse drug reaction
  *hyaluronic acid: CT, clinical trial
  *hyaluronic acid: AD, drug administration
  *hyaluronic acid: CM, drug comparison
  *hyaluronic acid: DO, drug dose
  *hyaluronic acid: DT, drug therapy
  *hyaluronic acid: PK, pharmacokinetics
  *hyaluronic acid: PD, pharmacology
  chondroitin sulfate: PR, pharmaceutics
hydroxypropylmethylcellulose: CM, drug comparison
ial: CM, drug comparison
ial: CT, clinical trial
ial: PD, pharmacology
ial: DT, drug therapy
ial: DO, drug dose
ial: PK, pharmacokinetics
ial: AD, drug administration
ial: AE, adverse drug reaction
lubricating agent: CT, clinical trial
lubricating agent: DT, drug therapy
lubricating agent: PK, pharmacokinetics
lubricating agent: PD, pharmacology
lubricating agent: AD, drug administration
lubricating agent: AE, adverse drug reaction
lubricating agent: DO, drug dose
lubricating agent: CM, drug comparison
methylprednisolone: CM, drug comparison
triamcinolone: CM, drug comparison
vitrax: DO, drug dose
vitrax: CM, drug comparison
vitrax: PK, pharmacokinetics
vitrax: PD, pharmacology
vitrax: AD, drug administration
vitrax: CT, clinical trial
vitrax: AE, adverse drug reaction
vitrax: DT, drug therapy
unclassified drug
(hyaluronic acid) 31799-91-4, 9004-61-9,
9067-32-7; (chondroitin sulfate)
9007-28-7, 9082-07-9; (hydroxypropylmethylcellulose)
9004-65-3; (methylprednisolone) 6923-42-8, 83-43-2; (triamcinolone)
```

RN

124-94-7

CN Healon; Viscoat; Amvisc; Hyalgan;
Healon gv; Ial; Vitrax

=> fil drugl

FILE 'DRUGLAUNCH' ENTERED AT 17:26:57 ON 15 MAR 2003 COPYRIGHT (C) 2003 IMSWORLD Publications Ltd

FILE COVERS 1982 TO 25 Feb 2003 (20030225/ED)

## => d all

L162 ANSWER 1 OF 1 DRUGLAUNCH COPYRIGHT 2003 IMSWORLD

AN 94:42952 DRUGLAUNCH

SO Drug Launches, (27 Jan 1992)

DN 1000793

CN Trade Name: VISCOAT
CO Manufacturer: Alcon
CO Corporation: Nestle

LNC Canada LND Oct 1991

CC N1B Local Anesthetics, except Dermatologicals

COMP Active Ingredient: sodium hyaluronate; chondroitin sulfate.

NC 2

TX Local anesthetic

DOSFM sol

LNP sol 0.5% 15 ml C\$106.00 (RPP)

=> fil wpix

FILE 'WPIX' ENTERED AT 17:46:09 ON 15 MAR 2003 COPYRIGHT (C) 2003 THOMSON DERWENT

FILE LAST UPDATED: 12 MAR 2003 <20030312/UP>
MOST RECENT DERWENT UPDATE: 200317 <200317/DW>
DERWENT WORLD PATENTS INDEX SUBSCRIBER FILE, COVERS 1963 TO DATE

- >>> SLART (Simultaneous Left and Right Truncation) is now
   available in the /ABEX field. An additional search field
   /BIX is also provided which comprises both /BI and /ABEX <<</pre>
- >>> PATENT IMAGES AVAILABLE FOR PRINT AND DISPLAY <<<

```
>>> FOR DETAILS OF THE PATENTS COVERED IN CURRENT UPDATES,
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 http://www.stn-international.de/training_center/patents/stn guide.pdf <<<
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    GUIDES, PLEASE VISIT:
    http://www.derwent.com/userguides/dwpi guide.html <<<
=> d all abeq tech abex tot
L188 ANSWER 1 OF 4 WPIX (C) 2003 THOMSON DERWENT
     2002-617109 [66]
                       WPIX
DNC C2002-174439
TΙ
     Chondroprotective/restorative composition useful for treating or
     preventing osteoarthritis and other joint diseases in mammals comprises
     hvaluronic acid or its salts.
     A96 B05 C03 D13
DC
     PIERCE, S W
IN
    (PIER-I) PIERCE S W
PA
CYC 1
                                                     A61K031-715
     US 2002068718 A1 20020606 (200266)* 14p
PI
ADT US 2002068718 A1 Provisional US 2000-237838P 20001003, US 2001-967977
     20011002
PRAI US 2000-237838P 20001003; US 2001-967977
                                                 20011002
     ICM A61K031-715
     ICS A61K031-70
     US2002068718 A UPAB: 20021014
AB
     NOVELTY - A chondroprotective/restorative composition comprises
     hyaluronic acid or its salts and optionally a
     pharmaceutical carrier.
          DETAILED DESCRIPTION - INDEPENDENT CLAIMS are also included for:
          (1) a method of treating or preventing osteoarthritis, joint
     effusion, joint inflammation and pain, synovitis, lameness, post operative
     arthroscopic surgery, deterioration of proper joint function, reduction or
     inhibition of metabolic activity of chondrocytes, activity of enzymes that
     degrade cartilage, reduction or inhibition of production of
     hyaluronic acid in mammals comprises oral administration
     of hyaluronic acid or its salt;
          (2) an animal feed having chondroprotective/restorative benefits
     comprising a nutritionally effective feed base selected from grains,
     proteins, and/or fats, and an hyaluronic acid or its
     salts; and
          (3) a therapeutic and chondroprotective/restorative composition
     comprising Hyaluronic acid or its salts, a therapeutic
     drug, and optionally a pharmaceutical carrier.
          ACTIVITY - Osteopathic; Antiarthritic; Anti-inflammatory; Analgesic.
          MECHANISM OF ACTION - None given.
          USE - Fort treating or preventing osteoarthritis, joint effusion,
     joint inflammation and pain, synovitis, lameness, post operative
     arthroscopic surgery, deterioration of proper joint function, reduction or
     inhibition of metabolic activity of chondrocytes, activity of enzymes that
     degrade cartilage, reduction or inhibition of production of
     hyaluronic acid in mammals. Hyaluronic
     acid, optionally in combination with glucosamine sulfate
     and/or chondroitin sulfate is useful in
     chondroprotective/restorative compositions. The composition is useful in
     an animal feed comprising a feed base selected from grains, proteins, fats
```

and mixtures of these. The animal feed further includes molasses. The animal feed is in the form of a paste and is a cat, dog or horse feed.

Dwg.0/0

```
CPI
FS
FA
    AB; DCN
    CPI: A12-V01; B01-B01; B01-B02; B01-C04; B02-Z; B03-L; B04-B01; B04-C02;
MC
          B05-A03; B05-B01P; B05-B02C; B05-C; B06-H; B07-H; B10-A04; B10-A06;
          B10-A07; B10-A08; B10-A10; B10-B02; B10-C04; B10-E04; B10-J02;
          B14-C01; B14-C03; B14-C09; B14-N01; C01-B01; C01-B02;
          C01-C04; C02-Z; C03-L; C04-B01C; C04-C02; C05-A03; C05-B01P;
          C05-B02C; C05-C01; C05-C08; C06-H; C07-H; C10-A04; C10-A06; C10-A07;
          C10-A08; C10-A10; C10-B02; C10-C04; C10-E04; C10-J02; C14-C01;
          C14-C03; C14-C09; C14-N01; D03-C; D03-F06; D03-G; D03-H01T2
TECH
                    UPTX: 20021014
    TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred drugs: The
    hyaluronic acid is uncrosslinked. The salt of
    hyaluronic acid is sodium hyaluronate
```

. The composition further comprises a supplement selected from vitamin A, D and E, ascorbic acid, biotin, pantothenic, choline, niacin, pyridoxine, riboflavin, thiamine, calcium, phosphorus, NaCl, copper, iron, manganese, iodine and combinations of these. Preferred composition: The composition further comprises glucosamine or its salt (preferably glucosamine sulfate) and/or chondroitin or its salt (preferably chondroitin sulfate). Preferably the composition further includes glucosamine sulfate and chondroitin sulfate. The composition further comprises a drug selected from acetaminophen, acetic acid, acetylsalicylic acid, buffered acetylsalicylic acid, albuterol, albuterol sulfate, ethanol isopropanol, allantoin, aloe, aluminum acetate, aluminum carbonate, aluminum chlorohydrate, aluminum hydroxide, alprozolam, amino acids, aminobenzoic acid, amoxicillin, ampicillin, amsacrine, amsalog, anethole, aspartame, atenolol, bacitracin, balsam peru, beclomethasone dipropionate, benzocaine, benzoic acid, benzophenones, benzoyl peroxide, biotin, bisacodyl, bornyl acetate, bromopheniramine maleate, buspirone, caffeine, calamine, calcium, calcium carbonate, calcium casinate, calcium hydroxide, camphor, captopril, cascara sagrada, castor oil, cephalosporins, cefaclor, cefadroxil, cephalexin, cetylalcohol, cetylpyridinium chloride, chelated minerals, chloramphenicol, chlorcyclizine hydrochloride. chlorhexidine gluconate, chloroxylenol, chloropentostatin, chlorpheniramine maleate, cholestyramine resin, choline bitartrate, cimetidine hydrochloride, cinnamedrine hydrochloride, citalopram, citric acid, clenbuterol, cocoa butter, cod liver oil, codeine and codeine phosphate, clonidine, clonidine hydrochloride, clorfibrate, ciprofloxacin HCl, cyanocobalamin, cyclizine hydrochloride, DMSO, danthron, dantrium, dexamethazone, dexbrompheniranime maleate, dextromethorphan hydrobromide, diazapam, dibucaine, diclofenac sodium, digoxin, diltiazem, dimethicone, dioxybenzone, diphenhydramine citrate, diphenhydramine hydrochloride, docusate calicum, docusate potassium, docusate sodium, doxycycline hyclate, doxylamine succinate, efaroxan, enalapril, enoxacin, erythromycin, estropipate, ethinyl estradiol, ephedrine, epinephrine bitartrate, erythropoietin, eucalyptol, ferrous fumarate, ferrous gluconate, ferrous sulfate, folic acid, fosphenytoin, flunixin meglumine, fluoxetine HCl, furosemide, gabapentan, gentamicin, gentocin sulfate, gemfibrozil, glipizide, glycerin, glyceryl stearate, griseofulvin, guaifenesin, hexylresorcinol, hydrochlorothiaxide, hydrocodone bitartrate, hydrocortisone, hydrocortisone acetate, 8-hydroxyquinoline sulfate , ibuprofen, indomethacin, inositol, insulin, iodine, ipecac, iron, isoxicam, isoxuprine, ketamine, ketofin, koalin, lactic acid, lanolin. lecithin, lidocaine, lidocaine hydrochloride, lifinopril, liotrix, lovastatin, MSM (methylsulfonylmethane), magnesium carbonate, magnesium hydroxide, magnesium salicylate, magnesium trisilocate, mefenamic acid, meclofenanic acid, meclofenamate sodium, medroxyprogesterone acetate, methenamine, mandelate, methocarbamol, menthol, meperidine hydrochloride, metaproterenol sulfate, methyl nicotinate, methyl salicylate, methylcellulose, methsuximide, metromidazole, metromidazole hydrochloride, metoprolol tartrate, miconazole nitrate, mineral oil,

minoxidil, morphine, naproxen, naproxen sodium, nifedipine, neomycin sulfate, neomycin-bacitracin, niacin, niacinamide, nicotine, nicotinamide, nitroglycerin, nonoxynol-9, norethindone, norethindone acetate, nystatin, octoxynol, octyl dimethyl PABA, octyl methoxycinnamate, omega-3 polyunsaturated fatty acids, omeprazole, oxolinic acid, oxybenzone, oxtriphylline, para-aminobenzoic acid (PABA), padimate O, paramethadione, penicillin, pentastatin, peppermint oil, pentaerythriol tetranitrate, pentobarbital sodium, pheniramine maleate, phenobarbital, phenol, phenolphthalein, phenybutazone, phenylephrine hydrochloride, phenylpropanolamine, phenylpropanolamine hydrochloride, phenytoin, pheneIzine sulfate, pirmenol, piroxicam, polymycin B sulfate, potassium chloride, potassium nitrate, prazepam, prednisone, prednisolone, procainamide hydrochloride, procaterol, propoxyphene, propoxyphene HCl, propoxyphene napsylate, pramiracetin, pramoxine, pramoxine hydrochloride, propronolol HCI, pseudoephedrine hydrochloride, pseudoephedrine sulfate, pyridoxine, quinapril, quinidine gluconate, quinestrol, ralitoline, ranitadine, resorcinol, riboflavin, salicylic acid. sesame oil, shark liver oil, simethicone, sodium bicarbonate, sodium citrate, sodium fluoride, sodium monofluorophosphate, sulfa-drugs, sulfanethoxazole, sulfur, tacrine, tacrine HCl, theophylline, terfenidine, thioperidone, trimetrexate, triazolam, timolol maleate, tretinoin, tetracycline hydrochloride, tolmetin, tolnaftate, triamcinolone, triclosan, triprolidine hydrochloride, undecylenic acid, vancomycin, verapamil HCl, vidaribine phosphate, vitamin A, vitamin B, vitamin C, vitamin D, vitamin E, vitamin K, witch hazel, xylometazoline hydrochloride, zinc, zinc sulfate and zinc undecylenate. The composition is in the form of a gel and comprises water and a sufficient amount of carboxymethylcellulose or its sodium salt.

**ABEX** 

ADMINISTRATION - The effective amount of hyaluronic acid is 10-2000 mg.

EXAMPLE - A paste was prepared from sodium hyaluronate (0.144 wt %), powdered sugar 10X (60.144 wt %), glycerine (0.7 wt %), xanthan gum (0.2 wt %), sodium benzoate (0.7 wt %), citric acid anhydrous (0.2 wt %), molasses (23.5 wt %) and water DI (14.4 wt %).

L188 ANSWER 2 OF 4 WPIX (C) 2003 THOMSON DERWENT

AN 2002-303912 [34] WPIX

DNC C2002-088337

TI Treatment of allergies, autoimmunity, adhesion cascade, metastatic or coronary cascade diseases e.g. arthritis comprises administration of at least one complex carbohydrate e.g. chondroitin sulfate

DC A96 B04 D21

IN BROWN, H G; BROWN, K K; COOPER, C A

PA (DERM-N) DERMAL RES LAB INC

CYC 96

PI WO 2002009728 A1 20020207 (200234)\* EN 61p A61K031-715

RW: AT BE CH CY DE DK EA ES FI FR GB GH GM GR IE IT KE LS LU MC MW MZ

NL OA PT SD SE SL SZ TR TZ UG ZW

W: AE AG AL AM AT AU AZ BA BB BG BR BY BZ CA CH CN CO CR CU CZ DE DK DM DZ EC EE ES FI GB GD GE GH GM HR HU ID IL IN IS JP KE KG KP KR KZ LC LK LR LS LT LU LV MA MD MG MK MN MW MX MZ NO NZ PL PT RO RU SD SE SG SI SK SL TJ TM TR TT TZ UA UG US UZ VN YU ZA ZW

AU 2001081368 A 20020213 (200238) A61K031-715

ADT WO 2002009728 A1 WO 2001-US41473 20010731; AU 2001081368 A AU 2001-81368 20010731

FDT AU 2001081368 A Based on WO 200209728

PRAI US 2000-222046P 20000731

IC ICM A61K031-715

AB WO 200209728 A UPAB: 20020528

NOVELTY - Treatment/prevention of diseases and conditions associated with allergies, autoimmunity, adhesion, metastatic or coronary cascades involves administration of at least one complex carbohydrate or a composition comprising at least one low purity or cosmetic grade complex carbohydrate and at least one transdermal or transmucosal carrier to deliver the complex carbohydrate into the blood stream.

DETAILED DESCRIPTION - Treatment or prevention of diseases associated with allergies, autoimmunity, adhesion cascade, metastatic cascade or coronary cascade involves: administration of at least one complex carbohydrate as sole active ingredient or a composition comprising at least one low purity or cosmetic grade complex carbohydrate as an active ingredient and at least one transdermal or transmucosal carrier to deliver the complex carbohydrate into the blood stream. The complex carbohydrate is oligosaccharide, sialylated oligosaccharide, polysaccharide or glycosaminoglycan.

INDEPENDENT CLAIMS are also included for the following:

- (1) interrupting the adhesion cascade by blocking the ability of leukocyte to bind to blood vessel walls, involving contacting the complex carbohydrate with receptor sites on leukocytes to inhibit the ability of the leukocyte to bind to the blood vessel walls to inhibit the motility to the site of trauma and thus reducing pain and swelling;
- (2) a bandage comprising either at least one complex carbohydrate and the carrier resulting in topical or mucosal delivery of the molecules, through the skin or mucous membranes of mammals and into the bloodstream or comprising only the complex carbohydrate added to it or imbedded in it. The bandage is applied onto an area requiring treatment; and
- (3) blocking the ability of tumor cells to tether to blood vessel walls by contacting the complex carbohydrates with receptor sites on tumor cells to inhibit the ability of the tumor cells to bind to the blood vessel walls and inhibit the tumor motility which, in turn, inhibits the potential for metastasis.

ACTIVITY - Immunosuppressive; Antiarthritic; Antirheumatic; Antiinflammatory; Antiulcer; Virucide; Antiallergic; Nootropic; Dermatological; Vasotropic; Vulnerary; Analgesic; Gynecological; Antiasthmatic; Antipruritic; Thrombolytic; Anticonvulsant; Tranquilizer; Neuroleptic; Neuroprotective; Antiparkinsonian; Cerebroprotective; Hypotensive; Cardiant; Anticoagulant; Anti-HIV; Antibacterial; Virucide; Antiseborrheic; Cytostatic; Antidiabetic; Antidepressant; Osteopathic.

MECHANISM OF ACTION - Macrophage inhibitor; T-cell inhibitor; Metastasis inhibitor; Tumor cell blocker; Amyloid plaque inhibitor; Leukocyte (CD44 and CD31) and RHAMM agonist; Leukocyte inhibitor.

USE - In the treatment of diseases associated with allergies, autoimmunity, adhesion cascade, metastatic cascade or coronary cascade e.g. arthritis, gastritis, colitis, stomach or intestinal ulcer, esophagitis, bronchitis, common cold, rhinitis, sore throat, tonsillitis, tendonitis, fibromyalgia, chronic fatigue syndrome, interstitial cystitis, polymyositis, autism, Lupus Erythematosis, headache, pancreatitis, anaphylaxis, vaginitis, hemorrhoids, sunburn, heat burn, temporomandibular joint (TMJ) condition, gingivitis, dental caries, dental pain, post surgical pain, menstrual pain, extremity cramp, pre and post partum pain, itching associated with allergies and hypersensitivity, asthma, emphysema, thrombosis, Attention Deficit Disorder, Attention Deficit Hyperactivity Disorder (ADHD), Turret's Syndrome, multiple sclerosis, Amyotrophic Lateral Sclerosis (ALS) or Lou Gehrig's Disease, Parkinson's Disease, Bell's Palsy, cerebral palsy, peripheral neuropathy, high blood pressure, heart disease, heart attack, vasculitis, stroke, increased degradation of spinal nerves post spinal cord injury, head and brain trauma post injury, encephalitis, epilepsy, Guillain-Barre syndrome, Human Immunodeficiency Virus infection, yeast infections, bacterial infections, viral infections, meningitis, peripheral neuropathy, Creuztfeldt-Jacob Disease, acne, cognitive disorder, adhesion formation post surgery or chemotherapy, scar formation post surgery, non-healing wounds, decubutis ulcers, irritation of nerve ganglion formation, Alzheimer's disease, human immunodeficiency

disease, ovarian cancer, lick granulomas, hot spots, eczema, wrinkling of skin, diabetes, scleroderma, skin problems, osteoarthritis, rashes, dementia, pain associated with cervical disc degeneration and hair loss; for inhibiting macrophages; for reducing scar tissue; as bandage (all claimed). Also in the treatment of rheumatoid arthritis, irritated or inflamed muscles, cramped muscles, inflamed tendons, inflamed nerves or nerve bundles (e.g. inflamed ganglion, trigger points), swollen and painful joints, inflamed bladder, bruised tissue, tired feet, open wounds, decubitis ulcers, inflamed stomach or intestinal lining, inflamed bronchi or esophagial lining, adhesions formed after surgery, trauma or chemotherapy, pain post surgery, dental work or injury, plaques formed on veins or arteries leading to heart disease and stroke, inflammation associated with Alzheimer's Disease, head or brain trauma, degration of the spinal cord post spinal cord injury, pain associated with insect bites or stings, tumor formation and tumor metastasis. The composition stimulates the healing of open wounds, increases cognitive function, thickens hair and fingernails, increases suppleness of skin.

ADVANTAGE - The method does not require pharmaceutical grade complex carbohydrates for the administration. As the composition is applied topically, orally, mucosally or parenterally the contaminants do not produce any adverse reactions.

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Dwg.0/2
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FS CPI

MC

FA AB; DCN

CPI: A12-V01; A12-V03A; B04-C02D; B04-C02E2; B14-A01; B14-A02; B14-A04; B14-C01; B14-C03; B14-C06; **B14-C09**; B14-E04; B14-E08; B14-E10; B14-F01B; B14-F02; B14-F04; B14-G02A; B14-G02D; B14-H01; B14-J01A2; B14-J01A3; B14-J01A4; B14-J01B4; B14-J05; B14-K01A; B14-L06; B14-N06A; B14-N07; B14-N13; B14-N14; B14-N16; B14-N17; B14-S01; B14-S04; D08-A05; D09-C

TECH UPTX: 20020528

TECHNOLOGY FOCUS - BIOLOGY - The reactive cells are white blood cells.

TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Method: The treatment step involves administering a mixture of high and low molecular weight sodium hyaluronate.

## ABEX

SPECIFIC COMPOUNDS - Sodium hyaluronate and chondroitin sulfate are specifically claimed as the complex carbohydrates.

ADMINISTRATION - The composition is administered orally, topically, mucosally or parenterally (claimed) on a repeated basis. The parenteral administration includes intramuscular, intravenous, subcutaneous, intradermal, intraperitoneal or injection routes. The composition is administered in a dosage of 0.000001 - 150 (preferably 0.001 - 100, especially 0.01 - 20) mg/kg.

EXAMPLE - A formulation (A) comprising hyaluronic acid powder made up to a 1% solution in deionized, distilled water was prepared. One half of the final 1% solution was removed and its molecular weight was broken down by alkaline hydrolysis. The pH was adjusted to 11 - 14 using 10N NaOH. The solution was then heated to a temperature of 37 - 50degreesC. When a molecular weight of 10000 - 50000 was obtained, the pH was adjusted back to neutral (pH = 6 - 7). The final mixture was prepared by combining 1 liter of the low molecular weight preparation with 1 liter of the original 1% solution of sodium hyaluronate. An

18 year old female suffering from chronic fibromyalgia localized in the face and neck for 5 years was used for study. Prior to use of (A), she had taken pain relievers, acupuncture and numerous other procedures to treat the condition. Nothing had provided substantial relief without severe side effects. She was given (A). She was asked to take (A) orally, holding the liquid in the mouth for several seconds to allow mucosal adsorption before swallowing it. She took 10 mg twice/day (0.2 mg/kg). She reported that

after only 1 day, her symptoms were improved. After one week of daily use, she reported no pain. She continued the treatment for 6 months and reported no return of fibromylagia.

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L188 ANSWER 3 OF 4 WPIX (C) 2003 THOMSON DERWENT
    1999-246164 [21]
                       WPIX
DNC C1999-072009
    Intra-articular composition useful for treatment of
TI
    arthropathy, comprises microcapsules comprising high-molecular substance
    and drug.
DC
    A11 A96 B05 B07 P32
    IMAMORI, K; ISHIGAKI, K; KASAI, S; OKADA, M; ONO, K; SUZUKI, M
ΙN
PΑ
     (SSSE) SSP CO LTD; (SSSE) SS PHARM CO; (SSSE) SS SEIYAKU KK
CYC
    30
                                                     A61K009-56
ΡI
                  A1 19990428 (199921) * EN
                                              28p
        R: AL AT BE CH CY DE DK ES FI FR GB GR IE IT LI LT LU LV MC MK NL PT
           RO SE SI
    CN 1215589
                  A 19990505 (199936)
                                                     A61K009-50
    CA 2251277
                  A1 19990427 (199941)
                                        EN-
                                                     A61K009-50
                                              12p
    JP 11222425 A 19990817 (199943)
                                                     A61K009-50
    KR 99037138
                  A 19990525 (200032)
                                                     A61K009-50
    US 6197326
                  B1 20010306 (200115)
                                                     A61F002-00
    US 6428804
                  B1 20020806 (200254)
                                                     A61F002-00
ADT EP 911025 A1 EP 1998-119414 19981014; CN 1215589 A CN 1998-124109
    19981027; CA 2251277 A1 CA 1998-2251277 19981020; JP 11222425 A JP
    1998-293385 19981015; KR 99037138 A KR 1998-43234 19981015; US 6197326 B1
    US 1998-172271 19981014; US 6428804 B1 Cont of US 1998-172271 19981014, US
    2000-706762 20001107
PRAI JP 1997-294009
                     19971027
    ICM A61F002-00; A61K009-50; A61K009-56
         A61F013-00; A61K009-08; A61K009-10; A61K009-107; A61K009-14;
         A61K009-52; A61K031-00; A61K031-57; A61K031-715; A61K031-73;
         A61K031-765; A61K031-78; A61K045-00; A61K047-30
AΒ
    EΡ
          911025 A UPAB: 20011203
    NOVELTY - A new intra-articular composition is useful
    for the treatment of arthropathy and comprises micro capsules comprising a
    high-molecular substance, which has biodegradability and biocompatibility,
    and a drug.
         USE - The composition is useful for the treatment of arthropathy and
    it is incorporated into the synovium or peripheral tissue within joints
    where it releases the drug over a sustained period of time.
         ADVANTAGE - The composition makes it possible to increase the
    concentration of a drug at a target area in a joint, to avoid general side
    effects, and to allow the sustained release of the drug, hence allowing
    the maintenance of the drug's efficacy over a long period of time.
         DESCRIPTION OF DRAWING(S) - The graph shows the relationship between
    time (days) and predonisolone released (%) from the predonisolone
    containing micro capsules (see example). It can be seen that the release
    is gradual over time in these compositions, e.g. only 70 % predonisolone
    released after 21 days, compared with 100 % predonisolone released after 1
    day with compositions comprising bulk powder predonisolone.
    Dwg.2/14
FS
    CPI GMPI
FΑ
    AB; GI; DCN
    CPI: A12-V01; B01-B02; B01-B03; B01-C02; B04-C03; B05-A01B; B05-A03B;
MC
         B05-B01J; B06-B02; B06-D01; B06-D02; B06-D09; B06-F02; B06-F03;
         B06-F04; B07-A02A; B07-A02B; B07-D04B; B07-D04C; B07-D09; B07-D13;
         B07-E01; B10-A07; B10-A10; B10-A23; B10-B02A; B10-B02D; B10-C03;
         B10-C04B; B10-C04C; B10-C04E; B10-F02; B12-M10A; B12-M11E
TECH
                    UPTX: 19990603
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TECHNOLOGY FOCUS - PHARMACEUTICALS - Preferred Micro capsules: The micro

capsules have an average particle size of 5-530 mum.

Preferred Drug: The drug is a steroidal agent, non steroidal

antiphlogistic, antirheumatic, immunoregulator, immunosuppressor or articular flow improver. The drug is at least one of dexamethasone, hydrocortisone, triamcinolone, beta-methosone, predonisolone, methylpredonisolone, halopredone, beclomethasone, deprodone, diclofenac, indomethacin, ibuprofen, ketoprofen, aspirin, diflunisal, fulfenamic acid, floctafenine, tolfenamic acid, sulindac, fenbufen, salicylic acid, acemetacin, proglumetacin, nabumetone, protizinic acid, thiaprofen, oxaprozin, loxoprofen, alminoprofen, zaltoprofen, flurbiprofen, flurbiprofen axetil, piroxicam, tenoxicam, ampiroxicam, meloxicam, D-penicillamine, bucillamine, gold sodium thiomalate, auranofin, lobenzarit, salazosulfapyridine, methotrexate, cyclophosphamide, azathioprine, mizoribine, cyclosporin and hyaluronic acid and salts thereof. The drug is present in the composition at 1-80 wt. %.

Preferred Composition: The composition is administered in a form suspended in a microcapsule dispersing medium containing at least one of hyaluronic acid, chondroitin sulfate and salts thereof.

TECHNOLOGY FOCUS - POLYMERS - Preferred High-Molecular Substance: The substance is at least one biodegradable, biocompatible high-molecular weight substance selected from homopolymers and copolymers of lactic acid, glycolic acid, caprolactone, valerolactone, butyrolactone, amino acids, alkyl cyanoacrylates and hydroxybutyrates; albumin; gelatin; starch; casein; and chitosan (preferably starch).

ABEX

ADMINSTRATION - The dosage form is an injection (claimed).

EXAMPLE - Lactic acid-glycolic acid copolymer (PLGA; comonomer molar ratio: 50/50, wt, average molecular wt. 124000) (2.5 g) and predonisolone (0.278 g) were dissolved in methylene chloride (347.2 g). The solution was sprayed in a spray granulator to give the desired micro capsules with a predonisolone content of 10.1 % and an average particle size of 5.61 mum. The micro capsules (0.5 mg) and bulk powder of predonisolone (0.5 mg) were placed in 10 ml aliquots of a phosphate buffer at pH 6.8, respectively, and the dissolved amount measured regularly. The results are shown in the graph. Generally it has been shown the release of predonisolone from the predonisolone-containing micro capsules was delayed compared with predonisolone from bulk powder of predonisolone.

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L188 ANSWER 4 OF 4 WPIX (C) 2003 THOMSON DERWENT
AN
     1994-036539 [05]
                        WPIX
DNC
    C1994-016777
     Compsns. for treating rheumatoid arthritis - contg. lipid-bound
TI
     glycosaminoglycan..
DC
     AOKI, S; IWASAKI, S; KIMATA, K; SUGIURA, N; SUZUKI, S
ΙN
PA
     (SEGK) SEIKAGAKU KOGYO CO LTD
CYC
    18
                   A1 19940202 (199405)* EN
                                              25p
PΙ
     EP 581282
                                                     A61K031-735
         R: AT BE CH DE DK ES FR GB GR IT LI LU NL SE
                                                     A61K031-725
     AU 9344314
                   A 19940203 (199411)
     JP 06072893
                   A 19940315 (199415)
                                              22p
                                                     A61K037-20
                                                     A61K031-725
     CA 2101482
                  A 19940131 (199416)
                                              18p
                                                     A61K037-22
     US 5470578
                  A 19951128 (199602)
     AU 668963
                   B 19960523 (199628)
                                                     A61K031-725
                                                     A61K031-735
     EP 581282
                   B1 19990512 (199923)
                                        ΕN
         R: AT BE CH DE DK ES FR GB GR IT LI LU NL SE
                                                     A61K031-735
                  E 19990617 (199930)
     DE 69324859
ADT EP 581282 A1 EP 1993-112169 19930729; AU 9344314 A AU 1993-44314 19930729;
     JP 06072893 A JP 1992-203558 19920730; CA 2101482 A CA 1993-2101482
     19930728; US 5470578 A US 1993-98936 19930729; AU 668963 B AU 1993-44314
     19930729; EP 581282 B1 EP 1993-112169 19930729; DE 69324859 E DE
     1993-624859 19930729, EP 1993-112169 19930729
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FDT AU 668963 B Previous Publ. AU 9344314; DE 69324859 E Based on EP 581282
PRAI JP 1992-203558
                      19920730
    4.Jnl.Ref; EP 466966; EP 493622
         A61K031-725; A61K031-735; A61K037-20; A61K037-22
IC
         A61K009-48; A61K031-715; C07H005-06
AB
     EΡ
           581282 A UPAB: 19940315
     Antirheumatic compsns. comprise a lipid-bound glycosaminoglycan (I) opt.
     in salt form, and a carrier. (I) are described in JA4-80201 and 4-80202.
          (I) comprises chondroitin sulphate, dermatan
     sulphate or hyaluronic acid bound to a
     glycerolipid, pref. a glycerophospholipid or glyceride, esp. phosphatidyl
     ethanolamine (PE) or phosphatidyl serine. (I) is prepd. by oxidising the
     reducing terminal of the glycosaminoglycan, lactonising the prod. and
     reacting the lactone with an NH2-contg. lipid to form an amide bond.
     Binding may also be via an aminoalkyl or ester bond. The compsns. are
     formulated as solns. for intra-articular injection.
          ADVANTAGE - The compsns. inhibit adhesion of inflammatory synovial
     membrane cells to joint cartilage tissue, alleviate inflammation of the
     synovial membrane, and have no toxicity or side effects.
     Dwq.1/5
     CPI
FS
     AB; DCN
FA
MC
     CPI: B04-C02V; B14-C06
          5470578 A UPAB: 19960115
     A method of treating rheumatism which comprises administering to mammals
     suffering from rheumatism a composition comprising between 0.1 to 80%
     lipid-bound glycosaminoglycan (gag) or a pharmaceutically acceptable salt
     thereof and a pharmaceutically acceptable carrier, wherein said
     composition is administered in a dose of 0.1 to 2,000 mg/adult once a day
     or within several weeks.
     Dwq.0/3
=> d his
     (FILE 'HOME' ENTERED AT 16:16:14 ON 15 MAR 2003)
                SET COST OFF
     FILE 'REGISTRY' ENTERED AT 16:16:25 ON 15 MAR 2003
                E CHONDROITIN SULFATE/CN
L1
              1 S E3
            136 S 9007-27-6/CRN AND 7664-93-9/CRN
L2
             11 S L2 AND 2/NC
L3
L4
              3 S L2 AND 9067-32-7/CRN
              2 S L4 NOT C4H6O2S
L5
L6
              1 S 9067-32-7
L7
              1 S 9004-61-9
              3 S 9004-61-9/CRN AND L2
\Gamma8
L9
              1 S L8 AND 3/NC
     FILE 'HCAPLUS' ENTERED AT 16:18:30 ON 15 MAR 2003
L10
              9 S L5 OR L9
           7390 S L1 OR L3
L11 ·
          11657 S CHONDROITIN(S) (SULFATE OR SULPHATE)
L12
            361 S CHONDROITINSULFATE OR CHONDROITINSULPHATE
L13
            200 S CHONDROITIN()(SULFURIC OR SULPHURIC)()ACID
L14
          11512 S CHONDROITIN(1W) (SULFATE OR SULPHATE OR (SULFURIC OR SULPHURIC
L15
           1628 S (CHONDROITINSULFURIC OR CHONDROITINSULPHURIC) () ACID
L16
          13151 S L11-L16
L17
L18
           1406 S L6
           1729 S (NA OR SODIUM) () (HYALURONATE OR HYALURON OR HYALURONIC ACID)
L19
L20
             83 S HEALON OR HYALGAN
```

25 S ARTZ OR FCH 200

L21

```
1862 S L18-L21
           9531 S L7
L23
          12860 S HYALURONATE OR HYALURON OR HYALURONIC ACID
L24
L25
           2450 S HYALURONAN
             65 S HYALURONAN (S) (NA OR SODIUM OR SODIUM SALT)
L26
           4638 S L17 AND L19-L26
L27
L28
            247 S L27 AND L22
     FILE 'REGISTRY' ENTERED AT 16:28:28 ON 15 MAR 2003
             11 S L1 OR L3
L29
                SEL RN
L30
             58 S E1-E11/CRN
             56 S L30 NOT L5, L9
L31
             38 S L31 NOT (MXS OR IDS)/CI
L32
L33
             18 S L31 NOT L32
L34
            262 S CHONDROITIN(L)SULFATE
             88 S L34 AND SALT
L35
L36
             63 S L35 NOT (MXS OR IDS)/CI
             31 S L36 NOT (COMPD OR WITH)
L37
     FILE 'HCAPLUS' ENTERED AT 16:31:37 ON 15 MAR 2003
L38
            484 S L37
L39
          13232 S L17, L38
            260 S L39 AND L22
L40
           4647 S L39 AND L23-L26
L41
            260 S L40, L41 AND L22
L42
            260 S L28, L42
L43
L44
             50 S L43 AND GEL?
L45
             24 S L43 AND VISCOELAST?
             1 S L43 AND INTRAARTICUL?
L46
              2 S L43 AND INTRA ARTICUL?
L47
             72 S L44, L45 NOT L46, L47
L48
             23 S L48 AND EYE?/CW
L49
                SEL DN AN 17
L50
              1 S L49 AND E12-E14
             49 S L48 NOT L49
L51
                SEL DN AN 5 24 38
              3 S E15-E23 AND L51
L52
                E CARTILAGE/CT
L53
          11561 S E3-E25
                E E3+ALL
          14712 S E7+NT
L54
                E JOINT/CT
L55
           4768 S E6-E28
                E E5+ALL
L56
           1255 S E2
                E JOINT/CT
                E E6+ALL
           8912 S E6, E5+NT
L57
L58
           2604 S E13+NT
                E OSTEOARTHRITIS/CT
           1853 S E3
L59
                E E3+ALL
L60
           2870 S E11, E12, E10+NT
              7 S CHONDRAL(L)LESION
L61
L62
             72 S ?CHONDRAL? (L) LESION
             17 S L43 AND L53-L62
L63
                SEL DN AN 3
              1 S L63 AND E1-E3
L64
                SEL DN AN L63 1 5 17
L65
              3 S E4-E12 AND L63
             15 S L10, L50, L52, L64, L65
L66
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E OCHOA/AU

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L67
              7 S E95
                E HERMIDA/AU
                E HUMBERTO/AU
                E ALCON/PA, CS
                E ALCOM/PA,CS
            786 S E3-E8
L68
            785 S ALCON?/PA,CS
L69
             12 S L67-L69 AND L43
L70
              1 S L67-L69 AND L10
L71
             2 S L67-L69 AND L66
L72
L73
             15 S L66, L71, L72
             10 S L70 NOT L73
L74
             15 S L73 AND L10-L28, L38-L74
L75
             10 S L74 AND L10-L28, L38-L75
L76
     FILE 'REGISTRY' ENTERED AT 16:53:16 ON 15 MAR 2003
              3 S L5 OR L9
L77
     FILE 'HCAPLUS' ENTERED AT 16:54:37 ON 15 MAR 2003
     FILE 'MEDLINE' ENTERED AT 16:55:10 ON 15 MAR 2003
L78
             56 S L77
             75 S VISCOAT
L79
             75 S L78, L79
L80
           3497 S L29 OR L37
L81
L82
           8170 S L12-L16
L83
           7265 S CHONDROITIN() (SULFATE OR SULPHATE)
                E CHONDROITIN SULFATE/CT
                E E18+ALL
           3467 S E7/CT, CN
L84
           8170 S L81-L83
L85
L86
           3497 S L1 OR L3
L87
           8170 S L85, L86
L88
            888 S L18-L21
L89
          10799 S L24-L26
L90
              0 S L6
           7519 S L7
L91
L92
          10849 S L88-L91
                E HYALURONIC ACID/CT
                E E3+AL
                E E3+ALL
          7519 S E20/CT, CN
L93
L94
          10849 S L92, L93
L95
           2288 S L87 AND L94
                E CARTILAGE/CT
                E E3+ALL
          42363 S E6+NT
L96
                E CARTILAGE/CT
                E E6+ALL
L97
           7773 S E6+NT
                E OSTEOARTHRITIS/CT
                E E3+ALL
L98
          21860 S E11+NT
                E JOINT/CT
                E E4+ALL
L99
         102887 S E4+NT
                E E4+ALL
                E JOINT DISEASE/CT
                E E5+ALL
L100
         165332 S E3+NT
                E KNEE/CT
           5874 S E3+NT
L101
                E SHOULDER/CT
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5032 S E3+NT
 L102
                  E E4+ALL
                  E SACROILIAC/CT
                  E E4+ALL
 L103
             1966 S E5+NT
                  E COXOFEMER/CT
                  E ANKLE/CT
 L104
             3343 S E3+NT
                  E ELBOW/CT
             3209 S E3+NT
 L105
             4957 S E4+NT
 L106
                  E WRIST/CT
             3805 S E3+NT
 L107
                  E INTERPHALANG/CT
                  E E5+ALL
 L108
              934 S E2+NT
 L109
             1092 S ?CHONDRAL? (L) LESION
 L110
              441 S L95 AND L96-L109
                  E INTRAARTICULAR/CT
                  E E4+ALL
 L111
                6 S L110 AND E2+NT
                  E E2+ALL
 L112
                0 S L80 AND L96-L109
 L113
                0 S L110 AND L80
                8 S L92 (L) TU/CT AND L110
 L114
                  SEL DN AN 3 6 8
· L115
               5 S L114 NOT E1-E9
 L116
               26 S L92 (L) (AD OR PD OR PK)/CT AND L110
 L117
               6 S L116 AND CHONDROITIN SULFATES (L) ME/CT
               20 S L116 NOT L117
 L118
               13 S L118 AND CHONDROITIN SULFATES/CT, CN
 L119
 L120
               7 S L118 NOT L119
               18 S L115, L119
 L121
                  SEL DN AN 5-7 13 15 18
 L122
               12 S L121 NOT E10-E27
               10 S L122 NOT EXOGENOUS/TI
 L123 .
                  SEL DN AN 2 8
 L124
                2 S L123 AND E28-E33
      FILE 'MEDLINE' ENTERED AT 17:13:33 ON 15 MAR 2003
      FILE 'EMBASE' ENTERED AT 17:13:39 ON 15 MAR 2003
 L125
             123 S L80
 L126
             4438 S L81
 L127
             7594 S L12-L16
 L128
             6445 S L83
                  E CHONDROITIN SULFATE/CT
                  E E3+ALL
 L129
             3628 S E1
 L130
             7594 S L126-L129
 L131
             7527 S L6 OR L7
 L132
            10048 S L19, L20, L21, L24, L25, L26
 L133
             7594 S L37 OR L130
            10048 S L131, L132
 L134
 L135
             2276 S L133 AND L134
             2294 S L125, L135
 L136
                  E CARTILAGE/CT
                  E E3+ALL
 L137
            22568 S E3+NT
                  E E16+ALL
 L138
            11877 S E4+NT
                  E JOINT/CT
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E E3+ALL

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95195 S E3+NT
L139
L140
          54903 S E8-E44
            936 S ?CHONDRAL? (L) LESION
L141
                E OSTEOARTHRITIS/CT
                E E3+ALL
          13443 S E26+NT
L142
                E JOINT DISEASE/CT
                E E4+ALL
                E E2+ALL
         163212 S E4+NT
L143
L144
            433 S L136 AND L137-L143
                E INTRAARTICULAR/CT
                E E6+ALL
L145
           3689 S E1+NT
L146
             40 S L144 AND L145
L147
             14 S L146 NOT AB/FA
              4 S L144 AND (NA OR SODIUM) () HYALUR?
L148
L149
            255 S L144 AND L6
              3 S L148 AND L149
L150
              2 S L125 AND L144
L151
                SEL DN AN 2
L152
              1 S L151 AND E1-E2
              1 S L152 AND L125-L152
L153
     FILE 'EMBASE' ENTERED AT 17:24:12 ON 15 MAR 2003
     FILE 'DRUGLAUNCH' ENTERED AT 17:24:20 ON 15 MAR 2003
                E HYALUR
            311 S E4-E6, E8
L154
L155
            331 S L19, L20, L24, L25, L26
            331 S L154, L155
L156
            184 S L12, L16, L83
L157
L158
             18 S L79
            146 S HYALURON? (L) SODIUM
L159
L160
             12 S L156, L159 AND L157
             23 S L158, L160
L161
L162
              1 S L161 NOT OPHTHALM?/CC
     FILE 'DRUGLAUNCH' ENTERED AT 17:26:57 ON 15 MAR 2003
     FILE 'WPIX' ENTERED AT 17:27:09 ON 15 MAR 2003
           2462 S L156, L159
L163
                E SODIUM HYALURON/DCN
                E E4+ALL
            147 S E2
L164
L165
              0 S R07175/PLE
L166
           2492 S L163, L164
           1556 S (C08B037-08 OR C08L005-08)/IC, ICM, ICS
L167
L168
           3738 S L163-L167
           2563 S L156/BIX OR L159/BIX
L169
L170
           3827 S L168, L169
           1161 S L83/BIX OR L12/BIX OR L13/BIX OR L14/BIX OR L15/BIX OR L16/BI
L171
                E CHONDROITIN/DCN
                E E4+ALL
            794 S E2 OR 1875/DRN OR R01875/PLE
L172
            292 S E4
L173
L174
             52 S E6
L175
            585 S L171-L174 AND L170
             14 S L79/BIX
L176
            489 S L175 AND HYALURON?/BIX AND CHONDROITIN?/BIX
L177
L178
             94 S L177 AND (GEL OR VISCOELAST? OR VISCO ELAST?)/BIX
L179
              8 S L177 AND (?INTRAARTICUL? OR ?INTRA ARTICUL?)/BIX
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SEL DN AN 5 8

L181 31 S P421/MO,M1,M2,M3,M4,M5,M6 AND L175 L182 5 S L175 AND (B14-C09A OR C14-C09A)/MC	
T.182 5 S T.175 AND (B14-C09A OR C14-C09A)/MC	
HIGE S DE LET STATE (DIT OF OIT OF OIT) / HO	
L183 27 S L175 AND (B14-C09? OR C14-C09?)/MC	
L184 36 S L181-L183	
L185 3 S A61P019-02/IC, ICM, ICS, ICA, ICI AND L175	5
L186 34 S L184 NOT L185	
SEL DN AN 3 8	
L187 2 S E5-E8	
L188 4 S L180, L187 AND L163-L187	

FILE 'WPIX' ENTERED AT 17:46:09 ON 15 MAR 2003